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Leu Val Asn Ile Phe Trp Phe Tyr Arg Glu Ala Phe Leu Val Pro Ala -40 -35 -30	
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Tyr Thr Glu Gln Ser Gln Ile Lys Gly Tyr Val Trp Arg Ser Ala Val -25 -20 -15	
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Gly Phe Leu Phe Trp Val Ile Val Leu Thr Ser Trp Ile Thr Ile Phe -10 -5 1 5	
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Gln Ile Tyr Arg Pro Arg Trp Gly Ala Leu Gly Asp Tyr Leu Ser Phe	
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Thr Ile Pro Leu Gly Thr Pro	
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ggg Gly	gcc Ala	cag Gln	cct Pro	caa Gln 1	cag Gln	gag Glu	cca Pro	ctg Leu 5	gcc Ala	ctg Leu	gtc Val	ttc Phe	cgc Arg 10	ttc Phe	ggc Gly	241
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	cac His 30															337
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tgg Trp	cag Gln	ccg Pro	ggc	cgc Arg 65	61 y 999	ccc Pro	tgt Cys	gtc Val	ttg Leu 70	cag Gln	gag Glu	tac Tyr	cag Gln	cag Gln 75	ttc Phe	433
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	gcc Ala		Asp									tga	ttgt	gta		767
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							cgc Arg							19	93
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							ccc Pro							28	39
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							gtc Val							43	33
							aag Lys 85							4.8	31
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	_			_	_	_	ctg Leu	_			_	_	 -	81	17

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ctc cag cag gac cca gaa gct ccc aca gtg ccc aag aag ggg agg agg Leu Gln Gln Asp Pro Glu Ala Pro Thr Val Pro Lys Lys Gly Arg Arg 240 245 250	961
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gct gac atc cca tcc ttg gaa cca gag ggg acc tca gcc tct Ala Asp Ile Pro Ser Leu Glu Pro Glu Gly Thr Ser Ala Ser 270 275 280	1051.
tageaggagg eteteetige tigeacteac cetteetat tgtetigece tgeatetggg	1111 1171
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Cys	Phe	Val	Asp	Ile	Arg 50	Arg	Phe	Gly	Arg	Trp 55	Asp	Leu	Gly	Gly	Lys 60	
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	Gln															
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	Glu															
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	Cys	95					100					105				
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-	Leu 110					115					120					
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                                                                      104
Cys Ala Val Val Leu Ala Gln Tyr Leu Trp Phe His Arg Arg Ser Leu
        -75
                            -70
cca ggc aag gcc atc tta gag att gga gca gga gtg agc ctt cca gga
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Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala Gly Val Ser Leu Pro Gly
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                                            -50
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Ile Leu Thr Ala Lys Cys Gly Ala Glu Val Ile Leu Ser Asp Ser Ser
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Glu Leu Pro His Cys Leu Glu Val Cys Arg Gln Ser Cys Gln Met Asn
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Asn Leu Pro His Leu Gln Val Val Gly Leu Thr Trp Gly His Ile Ser
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Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp Ile Ile Leu Ala Ser Asp
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Phe Leu Met His Lys Asn Pro Lys Val Gln Leu Trp Ser Thr Tyr Gln
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Met Lys Cys Val His Ile Pro Leu Glu Ser Phe Asp Ala Asp Lys Glu
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gat ata gca gaa tot acc ott cca gga aga cat aca gtt gaa atg otg
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WO 99/31236 -53- PCT/IB98/02122

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Val	Leu	His	Val	Ser	Pro	Glu	Glu	Pro	Ara	Thr	Len	775 T	Thr	Dro	λla	, 55
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~~~	ctc	200	cct		~+~		~~+	~~+							_ •	
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g at	gaato g gg	gac t	gt gt	t tt	c ca	ag ag	gc ac	a gt	a ga 11 As	c aa p Ly	a to	rt at	a tt	c aa e Ly	g ata 's Ile	
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IJ	le Gl	n Va	l Leu	Lys	Met	Leu	Pro	Arg	Glu	Lys	Leu	Arg	Arg	Arg	Glu	
		20					25					30				
			a caa													203
G.			s Gln	Ile	Asn		Ъуs	Lys	Glu	Arg		Lys	Tyr	Glu	Thr	
	35					40					45					222
	-		a aga	-											-	233
P)		.9 ту	s Arg	GIU	_	гЛг	nys	ьys	пÀг							
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PCT/IB98/02122

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                                                                      112
                         Met Val Ala Leu Asn Leu Ile Leu Val Pro
                                         -10
tgc tgc gct gct tgg tgt gac cca cgg agg atc cac tcc cag gat gac
                                                                      160
Cys Cys Ala Ala Trp Cys Asp Pro Arg Ile His Ser Gln Asp Asp
gtg ccc cgt agc tct gct gct gat act ggg tct gcg atg cag cgg cgt
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Val Pro Arg Ser Ser Ala Ala Asp Thr Gly Ser Ala Met Gln Arg Arg
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gag gcc tgg gct ggt tgg aga agg tca caa ccc ttc tct gtt ggt ctg
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Glu Ala Trp Ala Gly Trp Arg Arg Ser Gln Pro Phe Ser Val Gly Leu
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cct tct gct gaa aga ctc gag aac caa cca ggg aag ctg tcc tgg agg
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Pro Ser Ala Glu Arg Leu Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg
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tcc ctg gtc gga gag gga tat aga atc tgt gac ctc tgacaactgt
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Ser Leu Val Gly Glu Gly Tyr Arg Ile Cys Asp Leu
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gagtacaccc tttccaggaa taatgttttg ggaaacactg aaatgaaatc ttcccagtat
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cca agg ggc ctg ggt gct ggg gag ggg tca ggt agt cca gtg cgt cca Pro Arg Gly Leu Gly Ala Gly Glu Gly Ser Gly Ser Pro Val Arg Pro -35 -30 -25	151
cct gta tcc acc tgg ggc cct agc tgg gcc cag ctc ctg gac agt gtc Pro Val Ser Thr Trp Glŷ Pro Ser Trp Ala Gln Leu Leu Asp Ser Val -20 -15 -10	199
cta tgg ctg ggg gca cta gga ctg aca atc cag gca gtc ttt tcc acc Leu Trp Leu Gly Ala Leu Gly Leu Thr Ile Gln Ala Val Phe Ser Thr -5 1 5	247
act ggc cca gcc ctg ctg ctg ctt ctg gtc agc ttc ctc acc ttt gac Thr Gly Pro Ala Leu Leu Leu Leu Val Ser Phe Leu Thr Phe Asp 10 20 25	295
ctg ctc cat agg ccc gca ggt cac act ctg cca cag cgc aaa ctt ctc Leu Leu His Arg Pro Ala Gly His Thr Leu Pro Gln Arg Lys Leu Leu 30 35 40	343
acc agg ggc cag agt cag ggg gcc ggt gaa ggt cct gga cag cag gag Thr Arg Gly Gln Ser Gln Gly Ala Gly Glu Gly Pro Gly Gln Gln Glu 45 50 55	391
gct cta ctc ctg caa atg ggt aca gtc tca gga caa ctt agc ctc cag Ala Leu Leu Gln Met Gly Thr Val Ser Gly Gln Leu Ser Leu Gln 60 65 70	439
gac gca ctg ctg ctg ctc atg ggg ctg ggc ccg ctc ctg aga gcc Asp Ala Leu Leu Leu Leu Met Gly Leu Gly Pro Leu Leu Arg Ala 75 80 85	487
tgt ggc atg ccc ttg acc ctg ctt ggc ctg gct ttc tgc ctc cat cct Cys Gly Met Pro Leu Thr Leu Leu Gly Leu Ala Phe Cys Leu His Pro 90 95 100 105	535
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att gag ctg gaa cct ggg ctg agc tcc agt gct gcc tgt aat ggg aag  Ile Glu Leu Glu Pro Gly Leu Ser Ser Ala Ala Cys Asn Gly Lys -15 -5 1	153
gag atg tca cca acc agg caa ctc cgg agg tgc cct gga agt cat tgc Glu Met Ser Pro Thr Arg Gln Leu Arg Arg Cys Pro Gly Ser His Cys 5 10 15	201
ctg aca ata act gat gtt ccc gtc act gtt tat gca aca acg aga aag Leu Thr Ile Thr Asp Val Pro Val Thr Val Tyr Ala Thr Thr Arg Lys 20 25 30	249
cca cct gca caa agc agc aag gaa atg cat cct aaa tagcaccatt Pro Pro Ala Gln Ser Ser Lys Glu Met His Pro Lys 35 40 45	295
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Val Leu Cys Val Leu Leu Gln Ala Gln Gly Gly Tyr Arg Asp Lys	
atg agg atg cag aga atc aag gtc tgt gag aag cga ccc agc ata gat	147
Met Arg Met Gln Arg Ile Lys Val Cys Glu Lys Arg Pro Ser Ile Asp 10 15 20	
cta tgc atc cac cac tgt tca tgt ttc caa aag tgt gaa aca aat aag	195
Leu Cys Ile His His Cys Ser Cys Phe Gln Lys Cys Glu Thr Asn Lys 25 30 35	
ata tgc tgt tca gcc ttc tgt ggg aac att tgt atg agc atc cta	240
Ile Cys Cys Ser Ala Phe Cys Gly Asn Ile Cys Met Ser Ile Leu 40 45 50	

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Cag cag gtc cca gag aac tac ttc tat gtg cca gac ctg ggc cag gtg Gln Gln Val Pro Glu Asn Tyr Phe Tyr Val Pro Asp Leu Gly Gln Val 25 30 35 40	208
CCt gag att gat gtt cca tcc tac ctg cct gac ctg ccc ggc att gcc Pro Glu Ile Asp Val Pro Ser Tyr Leu Pro Asp Leu Pro Gly Ile Ala 45 50 55	256
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gcc cct ggc acc att cca gaa ctg ccc acc ttc cac act gag gta gcc Ala Pro Gly Thr Ile Pro Glu Leu Pro Thr Phe His Thr Glu Val Ala 75 80 85	352
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ttg atc ttc ggt ctc gga gca gtt tgg ggg ctt ggt gtg gac cct tcc Leu Ile Phe Gly Leu Gly Ala Val Trp Gly Leu Gly Val Asp Pro Ser -10 -5 1 5	160
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gga gtg cgt cag gtc ccg ggg ctg cat aat ggg acg aaa gcc ttt ctc Gly Val Arg Gln Val Pro Gly Leu His Asn Gly Thr Lys Ala Phe Leu 25 30 35	256
ttt caa gat act ccc aga agc ata aaa gca tcc act gct aca gct gaa Phe Gln Asp Thr Pro Arg Ser Ile Lys Ala Ser Thr Ala Thr Ala Glu 40 45 50	304
Cag ttt ttt cag aag ctg aga aat aaa cat gaa ttt act att ttg gtg Gln Phe Phe Gln Lys Leu Arg Asn Lys His Glu Phe Thr Ile Leu Val 55 60 65 70	352
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ttg cgt tac ttt gaa ttt tcc acg ttt tta ctt tct ttg tct ctc atc Leu Arg Tyr Phe Glu Phe Ser Thr Phe Leu Leu Ser Leu Ser Leu Ile -20 -15 -10	268

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ctc Leu att Ile caa Gln aac Asn cct Pro cca Pro 75 cgca	tcc Ser -5 cct Pro cta Leu acc Thr ttt Phe 60 ttt Phe	aac Asn ggc Gly cag Gln ccc Pro 45 cag Gln cta Leu	ctt Leu caa Gln cag Gln 30 ttc Phe caa Gln gct Ala	aac Asn tca Ser 15 cag Gln gcc Ala att Ile tgc Cys	caa Gln gct Ala cct Pro aac Asn ctt Leu	atc Ile I cag Gln tcg Ser caa Gln cct Pro 65 aacco	ggc Gly ctg Leu gct Ala cta Leu 50 ggc Gly	agc Ser ttt Phe aac Asn 35 aat Asn cgt Arg	agc Ser att Ile 20 aaa Lys cca Pro cag Gln	cac His tac Tyr aaa Lys acg Thr tcc ser tc ca	ctc Leu caa Gln gca Ala caa Gln ggc Gly 70 aagaa	Ger II -10 gac Asp atg Met gga Gly cat His 55 agc Ser aggag	cgc Arg tct ser aaa Lys 40 ctg Leu ctc Leu g aag	cca Pro tca Ser 25 atc Ile gca Ala acc Thr gctto	cac His 10 caa Gln cac His aaa Lys tca Ser	100 148 196 244 292 347 407 a 467
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Val G																
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His Phe Leu Ly	s Glu Thr Pr	o Gly Ser A	sn Gln Ile I	le Pro Ser Pro	
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cag atg ggt ga	160				9
Gln Met Gly As	n Arg Ser A	a Thr Leu I	vs Arg Gln S	er Leu Asp Gln	
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tgg																439
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Gly																
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gac	cat	180	aac	atc	aad	gat		att	tac	age	aac	190	atr	tat	aaa	775
Asp	_	_			_	_		-		_		-			-	.,,
-	195		2		3 -	200	5		- 2 -		205			-1-		
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Leu	Leu	Glu	Asn	Gly		Arg	Ala	Gly	Thr	-	Val	Leu	Glu	Tyr		
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Thr	PLO	ьец	GIII	230	neu	Pile	ATA	Mec	235	GIII	TYL	ser	GIII	240	GIY	
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Phe																
			245					250					255			
ctt				_	_	_			_		-					967
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Leu				-	_				-							1013
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vai	GIY	ser	ьeu	ьуs 310	Thr	ser	Ala	vai	315	ser	inr	ser	Int	320	sei	
caa	gag	cct	aaa		ctc	ctc	agt	gga		ααa	aaσ	ccc	ctc	-	ctc	1159
					Leu											
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Arg	Thr	-	Phe	Ser												
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		_			-							-			tccttc	
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ctt ccc aag cta tct tct tat tct gga tgg gtg gta gag cac gtc cta Leu Pro Lys Leu Ser Ser Tyr Ser Gly Trp Val Val Glu His Val Leu 45 50 55	196
ccc cat atg cag gag aac caa cct ctg tct gag act tcg cca tcc tct Pro His Met Gln Glu Asn Gln Pro Leu Ser Glu Thr Ser Pro Ser Ser 60 65 70	244
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Asn Lys Lys Gly Asn Val Leu Gln Leu Pro Asn Phe
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225

115

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Tyr Glu T	rp Tyr	Gln Glu	Tyr T	yr Val	Gly A	Ala .	Ala	His	Gly	Leu	Ala	
	130			135					140		•	
gga att t												633
Gly Ile T	yr Tyr	Tyr Leu			Ser 1	Leu	Gln		Ser	Gln	Gly	
	45			50				155				
aag tta c												681
Lys Leu H	is Ser	Leu Val		ro Ser	Val 1			Val	Cys	Gln	Leu	
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Lys Phe P	ro Ser	-	Tyr P	ro Pro			GIA	Asp	Asn	Arg		
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Ile Gln A												025
TIE GIN A	210	nys var	rne A	215	Olu .	<i></i>	- y -	Deu	220	1105	1110	
tat cag t		ast ata	atc t		tat (	aaa	tta	cta		aaσ	gga	873
Tyr Gln C	ve Ala	Asn Val	Tle T	ro Gln	Tvr	ggy Glv	Leu	Leu	Lvs	Lvs	Glv	• • •
	25	mp var		30	-/-	<b>U</b> _1		235	-,-	-7-	1	
tat ggg c		cac ggt			aat	acc	tat		ttc	ctq	aca	921
Tyr Gly L	eu Cvs	His Glv	Ser A	la Gly	Asn	Ala	Tyr	Ala	Phe	Leu	Thr	
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Phe Ala G	lu Trp	Cys Leu	Glu T	yr Gly	Glu l	His	Gly	Cys	Arg	Thr	Pro	
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gac acc c												1065
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Leu				~~+ ++.		~~~			·+~ ·		-at-aa-	1226
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Tyr Asn Cys Trp Glu Phe Pro Ser Met Leu Ala Leu Ser Gly Tyr I	le
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Leu Glu Leu Ser Arg Lys Ala Lys Leu Ala Ala Thr Ala Gly Ala Pr 85 90 95	ro
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His Ile Leu Ala Gly Ile Cys Gly Met Val Ala Ile Ser Trp Tyr A	.la
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Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly Ser  1 5 10 15  ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac  Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  20 25 30  ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc  Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro  35 40 45  gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag  Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln  50 55 60  tct cag gac cac agt gga atc ttt ggc ctg gta aca aac ctg gaa gag  Ser Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu  65 70 75  ctg gag gtg gac gat tgg gag ttc tgagcctctg caaactgtgc gcattctca  Leu Glu Val Asp Asp Trp Glu Phe  80 85  gccagggatg cagaggccac ccagaggccc ttcctgaggg ccggccacat tcccgccctc  ctgggcagat tgggtagaaa ggacattctt ccaggaaagt tgactgctgg ctgattgga	110 158 206 254 302 356 416 476
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Leu	Arg	Ala	Arg	Met	Ala	Gln	Arg	Leu	Asp	Gly	Ala	Arg	Phe	Arg	Tyr	
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		_	_	_					_	-	_	_	cag	_		877
Leu	Asn	Glu	Gln	Leu	Tyr	Ser	Gly	Pro	Ser	Ser	Ala	Ala	Gln	Arg	Leu	
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Phe	${\tt Gln}$	Glu	Asp	Pro	Glu	Ala	Phe	Leu	Leu	Tyr	His	Arg	Gly	Phe	Gln	
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Val	Pro		Glu	Asp	Glu	Ser		Asp	Val	Ala	Val		Cys	Leu	Ser	
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Leu	Gln	Leu	Gln	Pro	Cys		Tyr	Lys	Arg	Arg						
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			_					-	_		-				ctggc	
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                                                                   113
                         Met Thr Leu Leu Ser Phe Ala Ala Phe Thr
                                        -25
get get the tee gee etc ecc tgt tae tae ett ggg etg ttt eag egg
                                                                   161
Ala Ala Phe Ser Val Leu Pro Cys Tyr Tyr Leu Gly Leu Phe Gln Arg
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               -15
geg etc geg teg gte tte gae eea ett tge gtt tgt tea egt gtg etc
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                                              10
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Glu Asn Gly Pro Cys Pro Thr Lys Glu Ala Ala Gln Leu Val Gly Lys
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gcc tcg ctg gag atc gag tac caa gtt tta gat gga gca gga tta gat Ala Ser Leu Glu Ile Glu Tyr Gln Val Leu Asp Gly Ala Gly Leu Asp 35 40 45	244
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<223> Von Heijne matrix

## score 4.4 seq AVASSFFCASLFS/AV

<221> polyA\_site <222> 775..787

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gacageeegg acaaaggee acces		Gly Ala Val Val Ala	
	-20	-15	
		——	100
gtg gct tcc agt ttc ttt tgt	gca tot oto tto	tea get gtg cac aag	100
Val Ala Ser Ser Phe Phe Cys	Ala Ser Leu Phe	Ser Ala Val His Lys	
-10	-5	1	
ata gaa gag gga cat att ggg	qta tat tac aga	ggc ggt gcc ctg ctg	148
Ile Glu Glu Gly His Ile Gly	Val Tyr Tyr Arg	Gly Gly Ala Leu Leu	
5 10	15	20	
act tog acc ago ggo cot ggt			196
act teg ace age gge eet ggt	Dhe Hie Lee Met	Iou Dro Dhe Ile Thr	
Thr Ser Thr Ser Gly Pro Gly		heu Pro Phe 11e 1m1	
25	30	35	044
tca tat aag tct gtg cag acc	aca ctc cag aca	gat gag gtg aag aat	244
Ser Tyr Lys Ser Val Gln Thr	Thr Leu Gln Thr	Asp Glu Val Lys Asn	
40	45	50	
gta cct tgt ggg act agt ggt	ggt gtg atg atc	tac ttt gac aga att	292
Val Pro Cys Gly Thr Ser Gly	Glv Val Met Ile	Tyr Phe Asp Arg Ile	
55	60	65	
gaa gtg gtg aac ttc ctg gtc			340
Glu Val Val Asn Phe Leu Val	Dro yea yla Val	His Asp Tle Val Lvs	
	PIO ASII AIA VAI	80	
70 75		* -	388
aac tat act gct gac tat gac	aag gcc ctc atc	ttc aac aag atc cac	300
Asn Tyr Thr Ala Asp Tyr Asp			
85 90	95	100	
cac gaa ctg aac cag ttc tgc	agt gtg cac acg	ctt caa gag gtc tac	436
His Glu Leu Asn Gln Phe Cys	Ser Val His Thr	Leu Gln Glu Val Tyr	
105	110	115	
att gag ctg ttt gga ctg gaa		cag gaa tot toa	481
Ile Glu Leu Phe Gly Leu Glu	Ass Ass Bho Sor	Cln Clu Ser Ser	
	125	130	
120			541
taaaagggac cctgagcaag aacat	ttttc atagcagaca	ggaggactca tecacatege	
cagcaatcat aattaagcaa accgc	ctttt gcaccattta	agatttagga aatcatccaa	601
attactttta atgtttctgc agtag	aaaat gaatctaaat	tcattttata gggtttgtag	661
tcttttatct gttttggatt cactg	tgctt ttaagaaaaa	gttggtaaat ttgccgttga	721
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<210> 112

<211> 569

<212> DNA

<213> Homo sapiens

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<222> 26..562

<221> sig\_peptide

<222> 26..187

<223> Von Heijne matrix score 4.1

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											-50						
													gaa			10	0
Ser	Ser	Pro	Ser	Leu	Lys	Thr	Asp	Thr	Ser	Pro	Val	Leu	Glu.	Thr			
-45					-40					-35					-30		_
													gct			14	8
Gly	Thr	Val	Ala		Met	Ala	Ala	Thr		Ser	Ala	Arg	Ala		Ala		
				-25					-20					-15			
													agg			19	6
Ala	Val	Val		Ala	Ala	Ala	Arg		Gly	Ser	Glu	Ala	Arg	Val	Ser		
			-10					-5					1			2.4	
aag	gcc	gct	ttg	gct	acc	aag	ctg	ctg	tcc	ttg	agc	ggc	gtg	TTC	gcc	24	4
Lys		Ala	Leu	Ala	Thr		Leu	Leu	ser	Leu		GIY	Val	Pne	Ата		
	5					10					15				***	20	
gtg	cac	aag	ccc	aaa	999	CCC	act	tca	gcc	gag	ctg	ctg	aat	cgg	ttg	29	12
	His	Lys	Pro	Lys		Pro	Thr	Ser	Ala		Leu	Leu	Asn	Arg			
20					25					30			~~~		35	2.4	^
													gaa			34	U
гÀг	GIU	ьуs	ьеп		Ата	GIU	Ala	GIY		PIQ	ser	Pro	Glu	11p	1111		
				40				-++	45	~~+	~~~	~~~	2.C+		a a c	38	R
													act			30	
ьys	Arg	цуѕ	БУ В 55	GIII	1111	Ten	цуз	60	GIY	пть	Gry	GIY	Thr 65	Deu	ASP		
200	~~~	~~~		~~~	~++	cta	att		aas	att	aas	add	gga	aca	aaa	43	16
_	-	-	_		-								Gly				•
361	VIG	70	AL 9	Gry	VAI	שבע	75	vul	Cly	110	011	80	<b>4 1</b>		-,-		
ato	tta		agt	atσ	tta	tca		tac	aaα	agg	tat		gcc	att	qqa	4.8	34
													Ala				
	85		-		LCu	90	1		-,-	5	95				4		
gaa		aaa	aaa	act	act		aca	cta	gat	tct		aga	aag	qta	aca	53	32
													Lys				
100		1	-1-		105		<b>-</b>		2	110		2	•		115		
	gaa	aaa	cct	tac		atg	aac	ctc	atc		gtag					56	59
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<222> 4..279

<223> Von Heijne matrix score 6.8 seq AVMLYTWRSCSRA/IP

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	Met	TTE	-90	HIS	vaı	Thr	Leu	Glu -85	Asp	Ald	Leu	261	-80	Val	vah	
cta	ctt	gaa		ctt	ccc	ctc	ccc	gac	caq	cag	cca	tac		qaq	cct	96
Leu	Leu	Glu	Glu	Leu	Pro	Leu	Pro	Asp	Gln	Gln	Pro	Cys	Ile	Glu	Pro	
		-75					-70	•				-65				
cca	cct	tcc	tcc	atc	atg	tac	cag	gct	aac	ttt	gac	aca	aac	ttt	gag	144
Pro	Pro	Ser	Ser	Ile	Met	Tyr	Gln	Ala	Asn	Phe		Thr	Asn	Phe	Glu	
	-60					-55					-50				~a+	192
gac	agg	aat	gca	ttt	gtc	acg	ggc	att	gca	agg	Tac	att	gag	Gln	gc. Ala	192
_	Arg	ASN	Ala	Pne	-40	Int	GIY	Ile	WIG	-35	ıyı	116	GIG	GIII	-30	
-45 aca	atc	cac	tcc	agc		aat	gag	atg	cta	-	qaa	qqa	cat	gag		240
Thr	Val	His	Ser	Ser	Met	Asn	Glu	Met	Leu	Glu	Ğlu	Gly	His	Glu	Tyr	
·				-25					-20					-15		
gcg	gtc	atg	ctg	tac	acc	tgg	cgc	agc	tgt	tcc	cgg	gcc	att	ccc	cag	288
Ala	Val	Met	Leu	Tyr	Thr	Trp	Arg	Ser	Cys	Ser	Arg	Ala		Pro	Gln	
			-10					-5					1		2.52	226
gtg	aaa	tgc	aac	gag	cag	CCC	aac	cga	gta	gag	atc	Tac	gag	Tare	Thr	336
Val	Lys 5	Cys	Asn	GIU	Gin	10	ASI	Arg	Val	GIU	15	IYI	GIU	בעם	1111	
σta		ata	cta	gag	ccq		atc	acc	aaq	ctc		aaq	ttc	atg	tat	384
Val	Glu	Val	Leu	Glu	Pro	Glu	Val	Thr	Lys	Leu	Met	Lys	Phe	Met	Tyr	
20					25				-	30					35	
ttt	cag	cgc	aag	gcc	atc	gag	cgg	ttc	tgc	agc	gag	gtg	aag	cgg	ctg	432
Phe	Gln	Arg	Lys		Ile	Glu	Arg	Phe		Ser	Glu	Val	Lys		Leu	
				40					45			~~~	+	50	cta	480
tgc	cat	gcc	gag	cgc	agg	aag	gac	ttt Phe	gtc	Cer	Glu	Δla	Tvr	Leu	Leu	400
Cys	urs	Ald	55	Arg	Arg	Бур	Asp	60	Val	JCI	014	nıu	65			
acc	ctt	gac		ttc	atc	aac	atg	ttt	gct	gtc	ctg	gat	gag	cta	aag	528
Thr	Leu	Gly	Lys	Phe	Ile	Asn	Met	Phe	Ala	Val	Leu	qaA	Glu	Leu	Lys	
		70	_				75					80				
aac	atg	aag	tgc	agc	gtc	aag	aat	gac	cac	tcc	gcc	tac	aag	agg	gca	576
Asn		Lys	Cys	Ser	Val		Asn	Asp	His	Ser		Tyr	ьуѕ	Arg	Ala	
	85					90	~~~	~~+		C 3 C	95 tat	atc	cad	aaa	tca	624
gca	Cag	Dhe	Leu	Ara	Tare	Met	Δla	gat Asp	Pro	Gln	Ser	Ile	Gln	Glu	Ser	-
100		FIIC	шси	AT 9	105					110					115	
		ctt	tcc	atg			gcc	aac	cac	aac	agg	atc	acc	cag	tgt	672
Gln	Asn	Leu	Ser	Met	Phe	Leu	Ala	Asn	His	Asn	Arg	Ile	Thr	Gln	Cys	
				120					125					130		
ctc	cac	cag	caa	ctt	gaa	gtg	atc	cca	ggc	tat	gag	gag	ctg	ctg	gct	720
Leu	His	Gln			Glu	Val	Ile	Pro		Tyr	GIU	GIU			Ala	
			135			~-~	~~+	140				220	145		ctg	768
gac	acc	gtc	aac Nen	Tle	Cyc	. gcg . v∍l	Asn	י עמי ה שיי	Tvr	Glu	Asn	Lvs	Met	Tvr	Leu	, 53
ASP	110	150		. 110	Суз	Val	155		- ] -	014		160		- 2 -		
act	ccc			aaa	cat	atq		ctc	aag	gta	aaa	ctc	ccc			810
Thr	Pro	Ser	Glu	Lys	His	Met	Leu	Leu	Lys	Val	Lys	Leu	Pro	,		
	165					170					175					
tga	ggcc	gca	ссса	tgga	gc c	:tg <b>g</b> g	ctta	c cc	tctc	acct	tct	tctt	att	aaaa	atccgt	870
ttt	aaaa	aac	aaaa	aaaa	aa a	ıaa										893

<sup>&</sup>lt;210> 114

<sup>&</sup>lt;211> 1475

<sup>&</sup>lt;212> DNA

<sup>&</sup>lt;213> Homo sapiens

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<222> 1462..1475
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ggg cta gtg cga agc ccc tcg ctg gac cag atg ttc gac gcc gag
                                                                    153
Gly Leu Val Arg Ser Ser Pro Ser Leu Asp Gln Met Phe Asp Ala Glu
atc ctg ggc ttt tcc acc cct cca ggc cgg ctc tcc atg atg tcc ttc
                                                                    201
Ile Leu Gly Phe Ser Thr Pro Pro Gly Arg Leu Ser Met Met Ser Phe
            15
                               20
ate tte aac gee ete ace tgt gee etg gge ttg etg tae tte ate egg
                                                                    249
Ile Phe Asn Ala Leu Thr Cys Ala Leu Gly Leu Leu Tyr Phe Ile Arg
                           35
                                                                    297
cga gga aag cag tgt ctg gat ttc act gtc act gtc cat ttc ttt cac
Arg Gly Lys Gln Cys Leu Asp Phe Thr Val Thr Val His Phe Phe His
    45
                       50
                                           55
ctc ctg ggc tgc tgg ttc tac agc tcc cgt ttc ccc tcg gcg ctg acc
                                                                    345
Leu Leu Gly Cys Trp Phe Tyr Ser Ser Arg Phe Pro Ser Ala Leu Thr
                   65
                                       70
tgg tgg ctg gtc caa gcc gtg tgc att gca ctc atg gct gtc atc ggg
                                                                    393
Trp Trp Leu Val Gln Ala Val Cys Ile Ala Leu Met Ala Val Ile Gly
                80
                                   85
gag tac ctg tgc atg cgg acg gag ctc aag gag ata ccc ctc aac tca
                                                                    441
Glu Tyr Leu Cys Met Arg Thr Glu Leu Lys Glu Ile Pro Leu Asn Ser
            95
                               100 .
gcc cct aaa tcc aat gtc tagaatcagg ccctttggac atcccgctga
                                                                    489
Ala Pro Lys Ser Asn Val
cacttgggcc ccttaacacc ttgggctgct cagaccctcc agatgaggtc cagcccagat
                                                                    549
ctgagaggaa ccctggaaat gtgaagtctc tgttggtgtg ggagagatag tgagggcctg
                                                                    609
tcaaagaagg caggtagcag tcagcatgac agctgcaaga atgacctctg tctgttgaag
                                                                    669
cettggtate tgagaggtea ggaaggggae etetttgagg gtaataacat aattggaace
                                                                    729
atgccactct tgagccacaa tacctgtcac cagcctgttg ttttaagaga gaaaaaaaat
                                                                    789
                                                                    849
Caaggatate tgattggage aaaccaette tttagteate tgtettaeet eeetgggaca
gctgttacct ttgcagtgtt gccgaatcac agcagttacc tttgcaatgt tgccgaatca
cagcagttct gttggagaaa cgcttggttt ccggatccag agccacagaa agaaatgtag
gtgtgaagta ttaggctgct gtcagggaga ggatggcaga tggaggcatc aagcacaagg
                                                                   1029
aaaatgcaca acctgtgccc tgttatacac acgttcatgt gcgcccaaga acctatgact
                                                                   1089
ttetteeagt teettetace aggteeceat cetgetgeea geteteaaca tageaggeea
                                                                   1149
                                                                   1209
taggacccag agaagaatcc cagtgttgct caaagtctga ccatcataaa gacactgcct
gtettetagg aatgaceagg cacceagete ecactggact ceaatttttt tteetgeett
                                                                   1269
atttagaatt ctttggcggg aagggtatga tgggttccca gagacaagaa gcccaacctt
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	152
	200
	248
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51

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Arg	Arg	Gln	Arg	Leu	Ala -40	Glu	Leu	cag Gln	Ala	Lys -35	His	Gly	Asp	Pro	Gly -30	99
gat Asp	gcg Ala	gcc Ala	caa Gln	cag Gln -25	gaa Glu	gca Ala	aag Lys	cac His	agg Arg -20	gaa Glu	gca Ala	gaa Glu	atg Met	aga Arg -15	aac Asn	147
agt Ser	atc Ile	tta Leu	gcc Ala -10	caa Gln	gtt Val	ctg Leu	gat Asp	cag Gln -5	tcg Ser	gcc Ala	cgg Arg	gcc Ala	agg Arg 1	tta Leu	agt Ser	195
aac Asn	tta Leu 5	gca Ala	ctt	gta Val	aag Lys	cct Pro 10	gaa Glu	aaa Lys	act Thr	aaa Lys	gca Ala 15	gta Val	gag Glu	aat Asn	tac Tyr	243
ctt Leu 20	ata	cag Gln	atg Met	gca Ala	aga Arg 25	tat	gga Gly	caa Gln	cta Leu	agt Ser 30	gag Glu	aag Lys	gta Val	tca Ser	gaa Glu 35	291
caa	ggt Gly	tta Leu	ata Ile	gaa Glu 40	atc	ctt Leu	aaa Lys	aaa Lys	gta Val 45	agc Ser	caa Gln	caa Gln	aca Thr	gaa Glu 50	aag Lys	339
aca Thr	aca Thr	aca Thr	gtg Val 55	aaa	ttc Phe	aac Asn	aga Arg	aga Arg 60	aaa	gta Val	atg Met	gac Asp	tct Ser 65	gat Asp	gaa Glu	387
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gtc	tagg	aca	g													450
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	-	-	_sic .114													
	_	_	_sit													
	00> : gctg(	ctt a	Met (	gga d Gly H	cac o	egc t Arg I	tc o	Leu A	gc g Arg C	gc c	etc t Seu I	ta a Leu 1	Chr I	tg c Leu I	tg ct Leu Le	g 51 u
Pro	g ccg	g cca o Pro	a cc	cto	g tat ı Tyı	aco Thi	c cgg	g cad	c cgc	atg Met	g cto Lev 5	ggt Gly	cca Pro	gag Glu	g tcc 1 Ser	99
gte Va: 10	c cc	3 000	c cca	a aaa	a cga s Arq 15	a tco g Se:	c cgo	g Sei	c aaa r Lys	a cto Lei 20	c ato	g gca E Ala	a ccg	g cco Pro	c cga o Arg 25	147

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	Gly 999															133
gca Ala	ctg Leu	ctt Leu	cgc Arg 45	ctc Leu	ctg Leu	ccg Pro	gag Glu	tac Tyr 50	cgg Arg	gat Asp	gca Ala	gag Glu	att Ile 55	gtg Val	cgg Arg	243
acc Thr	cgg Arg	gat Asp	ccc	gaa Glu	aaa Lys	ctc Leu	Ala	tcc	tgt Cys	gac Asp	atc Ile	Val	gtg	gac Asp	gtg Val	291
ggg ggg	ggc Gly	60 gag Glu	tac Tyr	gac Asp	cct Pro	cgg Arg	65 aga Arg	cac His	cga Arg	tat Tyr	Asp	70 cat His	cac His	cag Gln	agg Arg	339
tct Ser	75 ttc Phe	aca Thr	gag Glu	acc Thr	atg Met	80 agc Ser	tcc Ser	ctg Leu	tcc Ser	cct Pro	85 ggg Gly	agg Arg	ccg Pro	tgg Trp	cag Gln	387
90 acc	aag	ctg	agc	agt	95 gcg	gga	ctc	atc	tat	100 ctg	cac	ttc	ggg	cac	105 aag	435
	Lys		,	110					115					120		483
Leu	Leu	Ala	Gln 125	Leu	Leu	Gly	Thr	Ser 130	Glu	Glu	Asp	Ser	Met 135	Val	Gly	
acc Thr	ctc Leu	tat Tyr 140	gac Asp	aag Lys	atg Met	tat Tyr	gag Glu 145	aac Asn	ttt Phe	gtg Val	gag Glu	gag Glu 150	gtg Val	gat Asp	gct Ala	531
gtg Val	gac Asp	aat	GJA aaa	atc Ile	tcc Ser	cag Gln 160	tgg	gca Ala	gag Glu	ggg ggg	gag Glu 165	cct Pro	cga Arg	tat Tyr	gca Ala	579
ctg Leu	155 acc Thr	act Thr	acc Thr	ctg Leu	agt Ser	gca	cga Arg	gtt Val	gct Ala	Arg	ctt	aat Asn	cct Pro	acc Thr	Trp	627
170 aac	cac	ccc	gac	caa	175 gac	act	gag	gca	999	180 ttc	aag	cgt	gca	atg	185 gat	675
	His gtt			190					195					200		723
Leu	Val	Gln	Glu 205	Glu	Phe	Leu	Gln	Arg 210	Leu	Asp	Phe	Tyr	Gln 215	His	Ser	
tgg Trp	ctg Leu	cca Pro 220	Ala	cgg Arg	gcc Ala	ttg Leu	gtg Val 225	Glu	gag Glu	gcc Ala	ctt Leu	gcc Ala 230	Gln	cga Arg	ttc Phe	771
	gtg	gac Asp	cca				att Ile	gtg				Lys				819
Pro	235 tgg Trp	aag	gag Glu	cat His	Leu	tac Tyr	cac	ctg Leu	gaa Glu	tct Ser 260	999 Gly	ctg	tcc Ser	cct Pro	cca Pro 265	867
250 gtg Val	gcc Ala	ato Ile	tto Phe	Phe	Val	ato	tac Tyr	act Thr	Asp	cag Gln	gct	gga Gly	cag Gln	tgg Trp 280	cga Arg	915
ata Ile	cag Gln	tgt Cys	Val	Pro	aag	gag Glu	ccc Pro	His	Ser	tto	caa Gln	ago Ser	cgg Arg 295	ctg Leu		963
cto Lev	g cca l Pro	Glu	Pro	ı tgg	cgg Arg	ggt Gly	Leu	Arg	gac	gag Glu	gcc Ala	Leu	gac Asp	cag	gtc Val	1011
agt Sei	: Gly	Ile	: cct	ggc Gly	tgo Cys	Ile	Phe	gto	cat His	gca Ala	Sex	Gly	tto	att	ggc	1059
Gly	/ His	: cgc	acc Thr	cga Arg	g Glu	Gly	gcc	ttg Lev	g ago	Met	Ala	: cgt	gcc Ala	acc Thr	ttg Leu	1107
	o c cag a Glr					cca						ata	aaac	ctto	345 :ca	1157

tctcaaaaaa aaaaaa

350

355

686

1173

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tee cae tee agg etg tee eec ega aag ace cae tta etg tae ate ete

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Ser His Ser Arg Leu Ser Pro Arg Lys Thr His Leu Leu Tyr Ile Leu 160 165 170 175 agg ccc tct cgg cag ctg taggggtggg gaccggggag cacctgcctg	734
Arg Pro Ser Arg Gln Leu 180	
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Cys Cys Lys Gly Gly Pro Asp Glu Asp Ala Val Glu Arg Gln Arg Arg	
-55 -50 -45 cag aag ttg ctt ctt gca caa ctg cat cac aga aaa agg gtg aag gca	151
Gln Lys Leu Leu Leu Ala Gln Leu His His Arg Lys Arg Val Lys Ala	
-40 -35 -30 -25	199
gct ggg cag atc cag gcc tgg tgg cgt ggg gtc ctg gtg cgc agg acc Ala Gly Gln Ile Gln Ala Trp Trp Arg Gly Val Leu Val Arg Arg Thr -20 -15 -10	
ctg ctg gtt gct gcc ctc agg gcc tgg atg att cag tgc tgg tgg agg Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln Cys Trp Trp Arg -5	247
acg ttg gtg cag aga cgg atc cgt cag cgg cgg cag gcc ctg ttg agg	295
Thr Leu Val Gln Arg Arg Ile Arg Gln Arg Arg Gln Ala Leu Leu Arg  10 15 20	343
gtc tac gtc atc cag gag cag gcg acg gtc aag ctc cag tcc tgc atc Val Tyr Val Ile Gln Glu Gln Ala Thr Val Lys Leu Gln Ser Cys Ile	243
25 30 35 40	
cgc atg tgg cag tgc cgg caa tgt tac cgc caa atg tgc aat gct ctc	391
Arg Met Trp Gln Cys Arg Gln Cys Tyr Arg Gln Met Cys Asn Ala Leu  45  50  55	439
tgc ttg ttc cag gtc cca gag agc agc ctt gcc ttc cag act gat ggc Cys Leu Phe Gln Val Pro Glu Ser Ser Leu Ala Phe Gln Thr Asp Gly	437
60 65 70	
ttt tta cag gtc caa tat gca atc cct tca aag cag cca gag ttc cac	487
Phe Leu Gln Val Gln Tyr Ala Ile Pro Ser Lys Gln Pro Glu Phe His 75 80 85	
att gaa atc cta tca atc tgaaaggcct ggggcatgga gaacaggctg	535
Ile Glu Ile Leu Ser Ile 90	
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                                                                       99
Asn Asp Ser Gln Leu Ser Ala Ser Phe Leu Gln Pro Ser Leu Gln Ala
                                 -25
aac tgt cct gct ttg gac cct gct gtg tca ctc tcc gca cca gcc ttt
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Asn Cys Pro Ala Leu Asp Pro Ala Val Ser Leu Ser Ala Pro Ala Phe
                                                 -5
                             -10
        -15
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Ala Ser Ala Leu Arg Ser Met Lys Ser Ser Gln Ala Ala Arg Lys Asp
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Asp Phe Leu Arg Ser Leu Ser Asp Gly Asp Ser Gly Thr Ser Glu His
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                                     25
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                                                                      291
 Ile Ser Ala Val Val Thr Ser Pro Arg Ile Ser Cys His Gly Ala Ala
                                 40
 att ecc acc gee egt gee etc tge eta gge tgt tee tge tge acc gaa
                                                                       339
 Ile Pro Thr Ala Arg Ala Leu Cys Leu Gly Cys Ser Cys Cys Thr Glu
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 cgc ctc ctc ctg cca ccg ccc tcc ctc ctt tct tta gaa gcc cct gcc
                                                                       387
 Arg Leu Leu Pro Pro Pro Ser Leu Leu Ser Leu Glu Ala Pro Ala
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                                                                       443
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 Ser Thr
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 atatattcag tttcaccaac aatgcatcaa gtacttttt ttttaagtaa agaaccgcag
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-96-

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cag	ggc	240 cac	caa	ctt	ttc	cat	245 atc	ttc	ttg	gtg	ctq	250 tqc	acq	ctq	gct	921
Gln	Gly 255	His	Gln	Leu	Phe	His 260	Ile	Phe	Leu	Val	Leu 265	Cys	Thr	Leu	Ala	
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270	neu	Giu	MIG	VAI	275	neu	wsb	TAT	GIU	280	Arg	Arg	PIO	TIE	285	
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GIU	Pro	Leu	HIS	7nr 290	His	Trp	Pro	His	Asn 295	Phe	Ser	Gly	Leu	Phe	Leu	
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Leu	Thr	Val	Gly 305	Ser	Ser	Ile	Leu	Thr	Ala	Phe	Leu	Leu	Ser 315	Gln	Leu	
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Val	Gln	Arg	Lys	Leu	Asp	Gln	Lys	Thr	Lys			_			_	
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						-		_	aaaa		330		-5- :		55	1213
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								Met	t Asp				a Ala	a Thi	Ala	
ttt	qta	att	act	tat	ata	ctt	agc	-20	o att	tee	acc	atc	-15	-	gca	102
									Ile							
ant	+	-10	~~~	~~~	<b>~</b> ~ ~	<b>++</b> -	-5	<b>+</b> - <b>-</b>	~~~	<b>_</b>		1				350
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5			_		10					15					20	
									atc Ile							198
	2211	Der	JEI	25	Tien	MO11	пåз	261	30	ттр	voħ	GIU	LIIG	35	261	
									gat							246
qea	GLU	Ala	Asp 40	GIu	ьуs	Tnr	ıyr	Asn 45	Asp	Ala	Leu	Phe	Arg 50	Tyr	Asn	

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aaa tgt gtg agt ttc aca cta act gag cag ttc atg gag aaa ttt gtt Lys Cys Val Ser Phe Thr Leu Thr Glu Gln Phe Met Glu Lys Phe Val 85 90 95 100	390
gat ccc gga aac cac aat agc ggg att gat ctc ctt agg acc tat ctt Asp Pro Gly Asn His Asn Ser Gly Ile Asp Leu Leu Arg Thr Tyr Leu 105 110 115	438
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ttt ggg gct ttg atc gga ctt tgt gct tgc att tgc cga agc tta tat Phe Gly Ala Leu Ile Gly Leu Cys Ala Cys Ile Cys Arg Ser Leu Tyr 135 140 145	534
ccc acc att gcc acg ggc att ctc cat ctc ctt gca gtg aca aag gag Pro Thr Ile Ala Thr Gly Ile Leu His Leu Leu Ala Val Thr Lys Glu 150 155 160	582
agc atg ctt cca gct gga gct gag tcc aag cac aca gcc act cct gca Ser Met Leu Pro Ala Gly Ala Glu Ser Lys His Thr Ala Thr Pro Ala 165 170 180	630
cac gca tgc gtg caa aca ggg aag ccc aag taggagaaga ggaaagaggt His Ala Cys Val Gln Thr Gly Lys Pro Lys 185 190	680
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catcttcttt tcttctccag tgaatatgat ctccaaaccc ttattttttc tttgaactgt aaaatttcca ctcatggacg atgcaaccaa cagatgcaat ctctgagaag atgaaaattg	860 920
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			Thr Ile Tyr Met Ala	
	-	_	cga agt cca gtt caa Arg Ser Pro Val Gln 20	150
Glu Asn Ser Ser A			gat gaa ttc att agt Asp Glu Phe Ile Ser 35	198
			cct ttt cga tac aat Pro Phe Arg Tyr Asn 50	246
			ata ccc aaa aac atg Ile Pro Lys Asn Met 65	294
			ttt gat gtg gtc aca Phe Asp Val Val Thr 80	342
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Asp Pro Gly Asn H			ctt agg acc tat ctt Leu Arg Thr Tyr Leu 115	438
			tta ggt ttg atg tgc Leu Gly Leu Met Cys 130	486
		s Ala Cys Ile	tgc cga agc tta tat Cys Arg Ser Leu Tyr 145	534
			gca gat acc atg ctg Ala Asp Thr Met Leu 160	582
			cagetgaaat eecaagetaa	642
			ggagccatcc tggatgtcca	702
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Pro	Суа	Ala	His 10	Glu	Ala	Leu	Leu	Asp 15	Glu	Asp	Thr	Leu	Phe 20	tgc Cys	Gln	146
Gly	Leu	Glu 25	Val	Phe	Tyr	Pro	Glu 30	Leu	Gly	Asn	Ile	Gly 35	Сув	aag Lys	Val	194
Val	Pro 40	Asp	Cys	Asn	Asn	Tyr 45	Arg	Gln	Lys	Ile	Thr 50	Ser	Trp	atg Met	Glu	242
Pro 55	Ile	Val	Lys	Phe	Pro 60	Gly	Ala	Val	Asp	Gly 65	Ala	Thr	Tyr	atc Ile	Leu 70	290
Val	Met	Val	Asp	Pro 75	Asp	Ala	Pro	Ser	Arg 80	Ala	Glu	Pro	Arg	cag Gln 85	Arg	338
Phe	Trp	Arg	His 90	Trp	Leu	Val	Thr	Asp 95	Ile	Lys	Gly	Ala	Asp 100	ctg Leu	Lys	386
Lys	Gly	Lys 105	Ile	Gln	Gly	Gln	Glu 110	Leu	Ser	Ala	Tyr	Gln 115	Ala	ccc Pro	Ser	434
Pro	Pro 120	Ala	His	Ser	Gly	Phe 125	His	Arg	Tyr	Gln	Phe 130	Phe	Val	tat Tyr	Leu	482
Cag Gln 135	gaa Glu	gga Gly	aag Lys	gtc Val	atc Ile 140	tct Ser	ctc Leu	ctt Leu	ccc Pro	aag Lys 145	gaa Glu	aac Asn	aaa Lys	act Thr	cga Arg 150	530
Gly	Ser	Trp	Lys	Met 155	Asp	Arg	Phe	Leu	Asn 160	Arg	Phe	His	Leu	ggc Gly 165	Glu	578
Pro	gaa Glu	gca Ala	agc Ser 170	acc Thr	cag Gln	ttc Phe	atg Met	acc Thr 175	cag Gln	aac Asn	tac Tyr	cag Gln	gac Asp 180	tca Ser	cca Pro	626
acc Thr	ctc Leu	Cag Gln 185	gct Ala	ccc Pro	aga Arg	gaa Glu	agg Arg 190	gcc Ala	agc Ser	gag Glu	ccc Pro	aag Lys 195	cac His	aaa Lys	aac Asn	674
cag Gln	gcg Ala 200	gag Glu	ata Ile	gct Ala	gcc Ala	tgc Cys 205	taga	tago	cg g	cttt	gcca		gggc	atgt		725
ggcc	ttcc	tg c	ccac	cacc	g ac a aa	gatg tcat	tggg ccaa	tat aaa	ggaa .aaaa	ccc aaa	cctc a	tgga	ta c	agaa	cccct	785 826

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                                       -35
ctg ccg gat gag gcc cgg agc ctg ccc ccg ccc aag ctg acc gac ccq
                                                                      157
Leu Pro Asp Glu Ala Arg Ser Leu Pro Pro Pro Lys Leu Thr Asp Pro
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                                     -20
cgg ctc ctc tac atc ggc ttc ttg ggc tac tgc tcc ggc ctg att gat
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Arg Leu Leu Tyr Ile Gly Phe Leu Gly Tyr Cys Ser Gly Leu Ile Asp
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                                -5
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Asn Leu Ile Arg Arg Arg Pro Ile Ala Thr Ala Gly Leu His Arg Gln
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                                            15
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Leu Leu Tyr Ile Thr Ala Phe Phe Leu Leu Asp Ile Ile Leu
                    25
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tccatccaat acgttgaagt cttcaaaatg cttgctccag tttcactgat acctgctgtt
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cctgaatttg atggaacatg tttcttatga cagttgaagc ttatgctaat ctgtatgttg
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gcccgtggtc ctcctaagtg tgagcttgcg gcggaccgag gcccacctgc ctccctgcct
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gettegecca ggaetegtga etgegteege agaagaaate acaacagege tggaattget
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tccaagttga aaacctttgg gtctttctat gcgaacggat tgaagaaacg caaaaagttt
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145

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Gln Ala Glu Arg Leu Phe Glu Asn Gln Leu Val Gly Pro Glu Ser Ile	
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Ala His Ile Gly Asp Val Met Phe Thr Gly Thr Ala Asp Gly Arg Val	•
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	•	_	115					120	_	-	-	Glu	125	•			
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Phe Arg Lys Asn Lys Thr Leu Gly Tyr Gly Val Pro Met Leu Leu Leu
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Lys Phe Asp Asp Trp Lys Asn Ile Arg Gly Pro Arg Pro Trp Glu Asp
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His Trp Tyr Ser Pro Pro Glu Arg Thr Gly Ile Ser Leu Ile Leu Thr

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<pre>&lt;222&gt; 838849  &lt;400&gt; 140 cccctatctc cagac agaatcaagg aggtt ctgtgttttg tgaag  ctc gtt atg gtg Leu Val Met Val</pre>	tgg atc aat Trp Ile Asn tac tgt gat Tyr Cys Asp aag ttg gaa Lys Leu Glu -70 gaa aat aca	ccctc ttcagtgd actat gac atg  Met  -115 gct ttt gtc a Ala Phe Val M  -100 ttg cta cat a Leu Leu His I  -85 cat ggg tcc t His Gly Ser I  ata tgg cct c Ile Trp Pro C	cct actatgttgc a cgc tgg tca tgt Arg Trp Ser Cys atg ctc acc acg Met Leu Thr Thr -95 aaa tca gct gct Lys Ser Ala Ala -80 cac agc aat gct Tyr Ser Asn Ala -65 caa ggg gtg ctg	tttctcccc 120 gag cac 174 Glu His -110 caa ctg 222 Gln Leu cac ctg 270 His Leu cca cag 318 Pro Gln gtg cgg 366
<pre>&lt;222&gt; 838849 &lt;400&gt; 140 cccctatctc cagac agaatcaagg aggtt ctgtgttttg tgaag  ctc gtt atg gtg Leu Val Met Val</pre>	tgg atc aat Trp Ile Asn tac tgt gat Tyr Cys Asp aag ttg gaa Lys Leu Glu -70 gaa aat aca Glu Asn Thr -55 tta tat aga	ccctc ttcagtgd actat gac atg  Met  -115 gct ttt gtc a Ala Phe Val M  -100 ttg cta cat a Leu Leu His I  -85 cat ggg tcc t His Gly Ser I  ata tgg cct c Ile Trp Pro C	cct actatgttgc a cgc tgg tca tgt Arg Trp Ser Cys atg ctc acc acg Met Leu Thr Thr -95 aaa tca gct gct Lys Ser Ala Ala -80 cac agc aat gct Tyr Ser Asn Ala -65 caa ggg gtg ctg Gln Gly Val Leu -50 cct tac aac gtg	### ### ##############################
<pre>&lt;222&gt; 838849 &lt;400&gt; 140 cccctatctc cagac agaatcaagg aggtt ctgtgttttg tgaag  ctc gtt atg gtg Leu Val Met Val</pre>	tgg atc aat Trp Ile Asn tac tgt gat Tyr Cys Asp aag ttg gaa Lys Leu Glu -70 gaa aat aca Glu Asn Thr -55 tta tat aga Leu Tyr Arg -40 tct cat gcc	ccctc ttcagtgd actat gac atg  Met  -115 gct ttt gtc a Ala Phe Val M  -100 ttg cta cat a Leu Leu His I -85 cat ggg tcc t His Gly Ser I  ata tgg cct c Ile Trp Pro G gcc atg ggg c Ala Met Gly I -35 cgc ttt tat t	cct actatgttgc a cgc tgg tca tgt Arg Trp Ser Cys atg ctc acc acg Met Leu Thr Thr -95 aaa tca gct gct Lys Ser Ala Ala -80 cac agc aat gct Tyr Ser Asn Ala -65 caa ggg gtg ctg Gln Gly Val Leu -50 cct tac aac gtg Pro Tyr Asn Val	### ### ##############################
<pre>&lt;222&gt; 838849 &lt;400&gt; 140 cccctatctc cagac agaatcaagg aggtt ctgtgttttg tgaag  ctc gtt atg gtg Leu Val Met Val</pre>	tgg atc aat Trp Ile Asn tac tgt gat Tyr Cys Asp aag ttg gaa Lys Leu Glu -70 gaa aat aca Glu Asn Thr -55 tta tat aga Leu Tyr Arg -40 tct cat gcc Ser His Ala	ccctc ttcagtgd actat gac atg  Met  -115 gct ttt gtc a Ala Phe Val M  -100 ttg cta cat a Leu Leu His I -85 cat ggg tcc t His Gly Ser I ata tgg cct c Ile Trp Pro G gcc atg ggg c Ala Met Gly I -35 cgc ttt tat t Arg Phe Tyr I -20 atc ctt att g	cct actatgttgc a cgc tgg tca tgt Arg Trp Ser Cys atg ctc acc acg Met Leu Thr Thr -95 aaa tca gct gct Lys Ser Ala Ala -80 cac agc aat gct Tyr Ser Asn Ala -65 caa ggg gtg ctg Gin Gly Val Leu -50 cct tac aac gtg Pro Tyr Asn Val ctc tta ttt cat Phe Leu Phe His -15 gag ggc ggt gtc	### State

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tcc atg													000
Ser Met	Ala Leu		Leu Phe	Cys	Asn	-	ıyr	vai	Leu	Pne	_	Leu	
		25				30					35		
ctc cgg	gac aga	ata ç	gta tta	ggc	agg	gca	tac	tcc	tac	cca	ctc	aac	654
Leu Arg	Asp Arg	Ile V	Val Leu	Gly	Arg	Ala	Tyr	Ser	Tyr	Pro	Leu	Asn	
_	40			_	45		_		_	50			
agt tat	gaa ctc	aaq	qca aac	taac	ctac	ct d	tcaa	caat	g ac	qqaq	raact	:	705
Ser Tvr	-		_	_	, 2				-		•		
	55	-,											
cagataaa		++a=+	a cotto			+ ~++	-ata	a+++	-++=+	- 22 2	+ = + +	+	765
_			-										
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Met Phe Thr Ser Thr Gly Ser Ser Gly Leu Tyr Lys Ala Pro Leu Ser
-30 -25 -20

Lys Ser Leu Leu Leu Val Pro Ser Ala Leu Ser Leu Leu Leu Ala Leu
-15 -5 1

Leu Leu Pro His Cys Gln Lys Pro Phe Val Tyr Asp Leu His Ala Val

Lys Asn Asp Phe Gln Ile Trp Arg Leu Ile Cys Gly Arg Ile Ile Cys 20 25 30

Leu Asp Leu Lys Asp Thr Phe Cys Ser Ser Leu Leu Ile Tyr Asn Phe 35 40 45

Arg Ilè Phe Glu Arg Arg Tyr Gly Ser Arg Lys Phe Ala Ser Phe Leu 50 60 65

Leu Gly Thr Trp Val Leu Ser Ala Leu Phe Asp Phe Leu Leu Ile Glu
70 75 80

Ala Met Gln Tyr Phe Phe Gly Ile Thr Ala Ala Ser Asn Leu Pro Ser 85 90 95

Gly Leu Ile Phe Cys Cys Ala Phe Cys Ser Glu Thr Lys Leu Phe Leu 100 105 110

Ser Arg Gln Ala Met Ala Glu Asn Phe Ser Ile 115 120

<210> 142

<211> 55

<212> PRT

<213> Homo sapiens

<400> 142

Met Ala Asp Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Gln Lys Arg 1 5 10 15

Met Tyr Tyr Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile Thr Phe

Phe Met Gly Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu Lys Gln 35 40 45

Lys Gln Lys Lys Arg Ser Asn

-115-

50

55

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<210> 143
<211> 67
<212> PRT
<213> Homo sapiens
<220>
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Met Ser Arg Asn Leu Arg Thr Ala Leu Ile Phe Gly Gly Phe Ile Ser
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Leu Ile Gly Ala Ala Phe Tyr Pro Ile Tyr Phe Arg Pro Leu Met Arg
Leu Glu Glu Tyr Lys Lys Glu Gln Ala Ile Asn Arg Ala Gly Ile Val
                            20
Gln Glu Asp Val Gln Pro Pro Gly Leu Lys Val Trp Ser Asp Pro Phe
Gly Arg Lys
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<210> 144
<211> 198
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -21..-1
<400> 144
Met Pro Val Pro Ala Leu Cys Leu Leu Trp Ala Leu Ala Met Val Thr
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Arg Pro Ala Ser Ala Ala Pro Met Gly Gly Pro Glu Leu Ala Gln His
Glu Glu Leu Thr Leu Leu Phe His Gly Thr Leu Gln Leu Gly Gln Ala
                                20
Leu Asn Gly Val Tyr Arg Thr Thr Glu Gly Trp Leu Thr Lys Ala Arg
                            35
Asn Ser Leu Gly Leu Tyr Gly Arg Thr Ile Glu Leu Leu Gly Gln Glu
                        50
Val Ser Arg Gly Arg Asp Ala Ala Gln Glu Leu Arg Ala Ser Leu Leu
                    65
Glu Thr Gln Met Glu Glu Asp Ile Leu Gln Leu Gln Ala Glu Ala Thr
                                     85
Ala Glu Val Leu Gly Glu Val Ala Gln Ala Gln Lys Val Leu Arg Asp
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140 145 150 155

Met Val Ala Gln Gln His Arg Leu Arg Gln Ile Gln Glu Arg Leu His
160 165 170

130

95 100 105 Ser Val Gln Arg Leu Glu Val Gln Leu Arg Ser Ala Trp Leu Gly Pro

Ala Tyr Arg Glu Phe Glu Val Leu Lys Ala His Ala Asp Lys Gln Ser

His Ile Leu Trp Ala Leu Thr Gly His Val Gln Arg Gln Arg Glu

Thr Ala Ala Leu Pro Ala

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175

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<211> 135
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -25..-1
<400> 145
Met Ser Leu Arg Asn Leu Trp Arg Asp Tyr Lys Val Leu Val Val Met
                                        -15
                   -20
Val Pro Leu Val Gly Leu Ile His Leu Gly Trp Tyr Arg Ile Lys Ser
Ser Pro Val Phe Gln Ile Pro Lys Asn Asp Asp Ile Pro Glu Gln Asp
                           15
Ser Leu Gly Leu Ser Asn Leu Gln Lys Ser Gln Ile Gln Gly Lys Xaa
Ala Gly Leu Gln Ser Ser Gly Lys Glu Ala Ala Leu Asn Leu Ser Phe
                    45
Ile Ser Lys Glu Glu Met Lys Asn Thr Ser Trp Ile Arg Lys Asn Trp
                                    65
Leu Leu Val Ala Gly Ile Ser Phe Ile Gly Asp His Leu Gly Thr Tyr
Phe Leu Gln Arg Ser Ala Lys Gln Ser Val Lys Phe Gln Ser Gln Ser
                            95
Lys Gln Lys Ser Ile Glu Glu
    105
<210> 146
<211> 255
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -70..-1
<400> 146
Met Gln Gln Lys Glu Gln Gln Phe Arg Glu Trp Phe Leu Lys Glu Phe
                    -65
                                        -60
Pro Gln Ile Arg Trp Lys Ile Gln Glu Ser Ile Glu Arg Leu Arg Val
                                    -45
Ile Ala Asn Glu Ile Glu Lys Val His Arg Gly Cys Val Ile Ala Asn
                                -30
Val Val Ser Gly Ser Thr Gly Ile Leu Ser Val Ile Gly Val Met Leu
```

Ala Pro Phe Thr Ala Gly Leu Ser Leu Ser Ile Thr Ala Ala Gly Val Gly Leu Gly Ile Ala Ser Ala Thr Ala Gly Ile Ala Ser Ser Ile Val 20 15 Glu Asn Thr Tyr Thr Arg Ser Ala Glu Leu Thr Ala Ser Arg Leu Thr 35 Ala Thr Ser Thr Asp Gln Leu Glu Ala Leu Arg Asp Ile Leu His Asp 50 Ile Thr Pro Asn Val Leu Ser Phe Ala Leu Asp Phe Asp Glu Ala Thr

-15

Lys Met Ile Ala Asn Asp Val His Thr Leu Arg Arg Ser Lys Ala Thr 80 85 Val Gly Arg Pro Leu Ile Ala Trp Arg Tyr Val Pro Ile Asn Val Val 100 95 Glu Thr Leu Arg Thr Arg Gly Ala Pro Thr Arg Ile Val Arg Lys Val 115 Ala Arg Asn Leu Gly Lys Ala Thr Ser Gly Val Leu Val Val Leu Asp 130 Val Val Asn Leu Val Gln Asp Ser Leu Asp Leu His Lys Gly Glu Lys 145 150 Ser Glu Ser Ala Glu Leu Leu Arg Gln Trp Ala Gln Glu Leu Glu Glu 160 165 Asn Leu Asn Glu Leu Thr His Ile His Gln Ser Leu Lys Ala Gly 175 180

<210> 147 <211> 59 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -49..-1

<210> 148 <211> 180 <212> PRT <213> Homo sapiens

 <400> 148

 Met Cys Ile Ser Gly Leu Cys Gln Ile Val Gly Cys Asp His Gln Leu

 1
 5
 10
 15

 Gly Ser Thr Val Lys Glu Asp Asn Cys Gly Val Cys Asn Gly Asp Gly
 20
 25
 30

 Ser Thr Cys Arg Leu Val Arg Gly Gln Tyr Lys Ser Gln Leu Ser Ala
 35
 40
 45

 Thr Lys Ser Asp Asp Thr Val Val Ala Ile Pro Tyr Gly Ser Arg His
 50
 60

 Ile Arg Leu Val Leu Lys Gly Pro Asp His Leu Tyr Leu Glu Thr Lys
 80

 Thr Leu Gln Gly Thr Lys Gly Glu Asn Ser Leu Ser Ser Thr Gly Thr
 80

 Phe Leu Val Asp Asn Ser Ser Val Asp Phe Gln Lys Phe Pro Asp Lys
 100

 Glu Ile Leu Arg Met Ala Gly Pro Leu Thr Ala Asp Phe Ile Val Lys
 120

 112 Arg Asn Ser Gly Ser Ala Asp Ser Thr Val Gln Phe Ile Phe Tyr

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135
Gln Pro Ile Ile His Arg Trp Arg Glu Thr Asp Phe Phe Pro Cys Ser
       150 155 160
Ala Thr Cys Gly Gly Gly Tyr Gln Leu Thr Ser Ala Glu Cys Tyr Asp
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                                                    175
Leu Arg Ser Asn
          180
<210> 149
<211> 162
<212> PRT
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<221> SIGNAL
<222> -23..-1
<400> 149
Met Gly Asp Lys Ile Trp Leu Pro Phe Pro Val Leu Leu Leu Ala Ala
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                             -15
Leu Pro Pro Val Leu Leu Pro Gly Ala Ala Gly Phe Thr Pro Ser Leu
                          1
Asp Ser Asp Phe Thr Phe Thr Leu Pro Ala Gly Gln Lys Glu Cys Phe
           15
                                    20
Tyr Gln Pro Met Pro Leu Lys Ala Ser Leu Glu Ile Glu Tyr Gln Val
                                35
Leu Asp Gly Ala Gly Leu Asp Ile Asp Phe His Leu Ala Ser Pro Glu
                             50
Gly Lys Thr Leu Val Phe Glu Gln Arg Lys Ser Asp Gly Val His Thr
                          65
Val Glu Thr Glu Val Gly Asp Tyr Met Phe Cys Phe Asp Asn Thr Phe
Ser Thr Ile Ser Glu Lys Val Ile Phe Phe Glu Leu Ile Pro Asp Asn
                  95
                                     100
Met Gly Glu Gln Ala Gln Glu Gln Glu Asp Trp Lys Lys Tyr Ile Thr
                                 115
               110
Gly Thr Asp Ile Leu Asp Met Lys Leu Glu Asp Ile Leu Val Ser Met
                             130
Val Phe
<210> 150
<211> 120
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -23..-1
<400> 150
Met Gly Asp Lys Ile Trp Leu Pro Phe Pro Val Leu Leu Leu Ala Ala
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Met Gly Asp Lys Ile Trp Leu Pro Phe Pro Val Leu Leu Leu Ala Ala -20 -15 -10

Leu Pro Pro Val Leu Leu Pro Gly Ala Ala Gly Phe Thr Pro Ser Leu -5 1 5

Asp Ser Asp Phe Thr Phe Thr Leu Pro Ala Gly Gln Lys Glu Cys Phe 10 15 20 25

Tyr Gln Pro Met Pro Leu Lys Ala Ser Leu Glu Ile Glu Tyr Gln Val 30 35 40

Leu Asp Gly Ala Gly Leu Asp Ile Asp Phe His Leu Ala Ser Pro Glu
45
Gly Lys Thr Leu Val Phe Glu Gln Arg Lys Ser Asp Gly Val His Thr
60
Cys Ile Arg Ser Lys Asn Gly Pro Gly Thr Ala Val His Ala Tyr Asn
75
Pro Ser Thr Phe Arg Gly Gln Val
90

<210> 151 <211> 7 <212> PRT <213> Homo sapiens <400> 151 Met Val Glu Met Thr Gly Val 1 5

<210> 152 <211> 199 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -42..-1

<400> 152 Met Asp Gly Gln Lys Lys Asn Trp Lys Asp Lys Val Val Asp Leu Leu -35 Tyr Trp Arg Asp Ile Lys Lys Thr Gly Val Val Phe Gly Ala Ser Leu -20 -15 Phe Leu Leu Ser Leu Thr Val Phe Ser Ile Val Ser Val Thr Ala - 5 Tyr Ile Ala Leu Ala Leu Leu Ser Val Thr Ile Ser Phe Arg Ile Tyr 15 Lys Gly Val Ile Gln Ala Ile Gln Lys Ser Asp Glu Gly His Pro Phe 35 30 Arg Ala Tyr Leu Glu Ser Glu Val Ala Ile Ser Glu Glu Leu Val Gln 45 Lys Tyr Ser Asn Ser Ala Leu Gly His Val Asn Cys Thr Ile Lys Glu 60 Leu Arg Arg Leu Phe Leu Val Asp Asp Leu Val Asp Ser Leu Lys Phe 80 Ala Val Leu Met Trp Val Phe Thr Tyr Val Gly Ala Leu Phe Asn Gly 95 Leu Thr Leu Leu Ile Leu Ala Leu Ile Ser Leu Phe Ser Val Pro Val 110 Ile Tyr Glu Arg His Gln Ala Gln Ile Asp His Tyr Leu Val Leu Ala 125 130 Asn Lys Asn Val Lys Asp Ala Met Ala Lys Ile Gln Ala Lys Ile Pro 145 140 Gly Leu Lys Arg Lys Ala Glu 155

<211> 43 <212> PRT <213> Homo sapiens <400> 153 Met Pro Phe Arg Met Ser Gly Tyr Ile Pro Phe Gly Thr Pro Ile Val 10 Ser Val Thr Phe Lys Gly Phe Pro Phe Leu Lys Asn Tyr Phe Lys Cys Leu Thr Leu Cys Tyr Cys Ser Arg Val Phe Asp <210> 154 <211> 50 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -37..-1 <400> 154 Met Glu Trp Ala Gly Lys Gln Arg Asp Phe Gln Val Arg Ala Ala Pro -30 -35 Gly Trp Asp His Leu Ala Ser Phe Pro Gly Pro Ser Leu Arg Leu Phe -10 -15 Ser Gly Ser Gln Ala Ser Val Cys Ser Leu Cys Ser Gly Phe Gly Ala Gln Glu <210> 155 <211> 153 <212> PRT <213> Homo sapiens <400> 155 Thr Val Pro Leu Leu Glu Pro Ala Asp His Ala Arg Gly Arg Ala 10 His Val His Leu Pro Glu Asn Val Arg Ser Gln Ser Pro Gly His Val 25 Arg Arg Gly Arg Ser Gly Ala Gln Val Leu Pro Thr Gly Pro Asp Glu 40 Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu 55 Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr 75 70 Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser 90 Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys 105 100 Trp Gly Glu Leu Leu Arg Thr Ile Pro Glu Ile Pro Pro Lys Arg Gly 120 Glu Leu Lys Thr Glu Leu Leu Gly Leu Lys Glu Arg Lys His Lys Pro 135

Gln Val Ser Gln Gln Glu Glu Leu Lys

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<210> 156
<211> 67
<212> PRT
<213> Homo sapiens
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Met Arg Gln Lys Arg Lys Gly Asp Leu Ser Pro Ala Lys Leu Met Met
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Leu Thr Ile Gly Asp Val Ile Lys Gln Leu Ile Glu Ala His Glu Gln
                                25
            20
Gly Lys Asp Ile Asp Leu Asn Lys Val Arg Thr Lys Thr Ala Ala Lys
                            40
                                                45
Tyr Gly Leu Ser Ala Gln Pro Arg Leu Val Asp Ile Ile Ala Ala Val
Pro Pro Glu
65
<210> 157
<211> 87
<212> PRT
<213> Homo sapiens
<400> 157
Met Asp Glu Leu Ser Glu Glu Asp Lys Leu Thr Val Ser Arg Ala Arg
                                     10
Lys Ile Gln Arg Phe Leu Ser Gln Pro Phe Gln Val Ala Glu Val Phe
                                 25
Thr Gly His Met Gly Lys Leu Val Pro Leu Lys Glu Thr Ile Lys Gly
Phe Gln Gln Ile Leu Ala Gly Glu Tyr Asp His Leu Pro Glu Gln Ala
                        55
Phe Tyr Met Val Gly Pro Ile Glu Glu Ala Val Ala Lys Ala Asp Lys
Leu Ala Glu Glu His Ser Ser
                85
<210> 158
<211> 250
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -85..-1
Met Ser Ala Glu Val Lys Val Thr Gly Gln Asn Gln Glu Gln Phe Leu
                                         -75
-85
                     -80
Leu Leu Ala Lys Ser Ala Lys Gly Ala Ala Leu Ala Thr Leu Ile His
                 -65
Gln Val Leu Glu Ala Pro Gly Val Tyr Val Phe Gly Glu Leu Leu Asp
                                 -45
Met Pro Asn Val Arg Glu Leu Xaa Ala Arg Asn Leu Pro Pro Leu Thr
                             -30
                                                 -25
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Glu Ala Gln Lys Asn Lys Leu Arg His Leu Ser Val Val Thr Leu Ala

Ala Lys Val Lys Cys Ile Pro Tyr Ala Val Leu Leu Glu Ala Leu Ala

-10

-15

Leu Arg Asn Val Arg Gln Leu Glu Asp Leu Val Ile Glu Ala Val Tyr 20 Ala Asp Val Leu Arg Gly Ser Leu Asp Gln Arg Asn Gln Arg Leu Glu Val Asp Tyr Ser Ile Gly Arg Asp Ile Gln Arg Gln Asp Leu Ser Ala 50 Ile Ala Arg Thr Leu Gln Glu Trp Cys Val Gly Cys Glu Val Val Leu Ser Gly Ile Glu Glu Gln Val Ser Arg Ala Asn Gln His Lys Glu Gln 85 Gln Leu Gly Leu Lys Gln Gln Ile Glu Ser Glu Val Ala Asn Leu Lys 100 Lys Thr Ile Lys Val Thr Thr Ala Ala Ala Ala Ala Thr Ser Gln 115 Asp Pro Glu Gln His Leu Thr Glu Leu Arg Glu Pro Ala Pro Gly Thr 130 Asn Gln Arg Gln Pro Ser Lys Lys Ala Ser Lys Gly Lys Gly Leu Arg 145 150 Gly Ser Ala Lys Ile Trp Ser Lys Ser Asn 160

<210> 159 <211> 24 <212> PRT <213> Homo sapiens

<210> 160 <211> 228 <212> PRT <213> Homo sapiens

Met Pro Thr Asn Cys Ala Ala Ala Gly Cys Ala Thr Thr Tyr Asn Lys His Ile Asn Ile Ser Phe His Arg Phe Pro Leu Asp Pro Lys Arg Arg 25 Lys Glu Trp Val Arg Leu Val Arg Arg Lys Asn Phe Val Pro Gly Lys 40 His Thr Phe Leu Cys Ser Lys His Phe Glu Ala Ser Cys Phe Asp Leu Thr Gly Gln Thr Arg Arg Leu Lys Met Asp Ala Val Pro Thr Ile Phe 75 70 Asp Phe Cys Thr His Ile Lys Ser Met Lys Leu Lys Ser Arg Asn Leu 90 Leu Lys Lys Asn Asn Ser Cys Ser Pro Ala Gly Pro Ser Ser Leu Lys Ser Asn Ile Ser Ser Gln Gln Val Leu Leu Glu His Ser Tyr Ala Phe 120 Arg Asn Pro Met Glu Ala Lys Lys Arg Ile Ile Lys Leu Glu Lys Glu 135 Ile Ala Ser Leu Arg Arg Lys Met Lys Thr Cys Leu Gln Lys Glu Arg

Ile Leu Pro Glu Lys Asn Cys Asn Ser Arg His Ala Gly Phe Val Gly

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Pro Ala Lys Leu Arg Gln

65

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<400> 163 Met Gln Asn Val Ile Asn Thr Val Lys Gly Lys Ala Leu Glu Val Ala -50 Glu Tyr Leu Thr Pro Val Leu Lys Glu Ser Lys Phe Arg Glu Thr Gly -35 Val Ile Thr Pro Glu Glu Phe Val Ala Ala Gly Asp His Leu Val His -20 His Cys Pro Thr Trp Gln Trp Ala Thr Gly Glu Glu Leu Lys Val Lys -5 Ala Tyr Leu Pro Thr Gly Lys Gln Phe Leu Val Thr Lys Asn Val Pro 15 Cys Tyr Lys Arg Cys Lys Gln Met Glu Tyr Ser Asp Glu Leu Glu Ala 30 Ile Ile Glu Glu Asp Asp Gly Asp Gly Gly Trp Val Asp Thr Tyr His 45 Asn Thr Gly Ile Thr Gly Ile Thr Glu Ala Val Lys Glu Ile Thr Leu 60 65 Glu Asn Lys Asp Asn Ile Arg Leu Gln Asp Cys Ser Ala Leu Cys Glu 75 80 Glu Glu Glu Asp Glu Asp Glu Gly Glu Ala Ala Asp Met Glu Glu Tyr 90 95 Glu Glu Ser Gly Leu Leu Glu Thr Asp Glu Ala Thr Leu Asp Thr Arg 110 Lys Ile Val Glu Ala Cys Lys Ala Lys Thr Asp Ala Gly Gly Glu Asp 125 130 Ala Ile Leu Gln Thr Arg Thr Tyr Asp Leu Tyr Ile Thr Tyr Asp Lys 140 145 Tyr Tyr Gln Thr Pro Arg Leu Trp Leu Phe Gly Tyr Asp Glu Gln Arg Gln Pro Leu Thr Val Glu His Met Tyr Glu Asp Ile Ser Gln Asp His 175 Val Lys Lys Thr Val Thr Ile Glu Asn His Pro His Leu Pro Pro 190 Pro Met Cys Ser Val His Pro Cys Arg His Ala Glu Val Met Lys Lys 205 210 Ile Ile Glu Thr Val Ala Glu Gly Gly Glu Leu Gly Val His Met 220 225 Tyr Leu Leu Ile Phe Leu Lys Phe Val Gln Ala Val Ile Pro Thr Ile

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<210> 164
<211> 89
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -80..-1
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Glu Tyr Asp Tyr Thr Arg His Phe Thr Met

250

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-25
       -30
Gln Leu Gly Arg Gly Leu Leu Ser Ala Cys Ala Pro Trp Gly Asp Gly
           -10
Ser Thr Gln Pro Val Pro Leu Cys Ser
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<210> 165
<211> 98
<212> PRT
<213> Homo sapiens
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<221> SIGNAL
<222> -15..-1
<400> 165
Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu Ala Val Leu Ala Trp
                   -10
                                       -5
Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg Met Lys Ser Arg Glu
Gln Gly Gly Arg Leu Gly Ala Glu Ser Arg Thr Leu Leu Val Ile Ala
                            25
His Pro Asp Asp Glu Ala Met Phe Phe Ala Pro Thr Val Leu Gly Leu
                        40
Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys Phe Ser Ala Val Phe
                    55
Arg Arg Glu Leu Ser Glu Tyr Thr Glu Gly Leu Thr Ser Glu Pro Leu
                70
Thr Ala
<210> 166
<211> 92
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<222> -36..-1
<400> 166
Met Leu Val Thr Gln Gly Leu Val Tyr Gln Gly Tyr Leu Ala Ala Asn
                        -30
                                            -25
Ser Arg Phe Gly Ser Leu Pro Lys Val Ala Leu Ala Gly Leu Leu Gly
                                        -10
                    -15
Phe Gly Leu Gly Lys Val Ser Tyr Ile Gly Val Cys Gln Ser Lys Phe
His Phe Phe Glu Asp Gln Leu Arg Gly Ala Gly Phe Gly Pro Gln His
                                                25
                            20
Asn Arg His Cys Leu Leu Thr Cys Glu Glu Cys Lys Ile Lys His Gly
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30 35 40 Leu Ser Glu Lys Gly Asp Ser Gln Pro Ser Ala Ser 45 50 55

<210> 167 <211> 351

<212> PRT

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<213> Homo sapiens

<220>
<221> SIGNAL
<222> -16..-1

<400> 167 Met Val Pro Phe Ile Tyr Leu Gln Ala His Phe Thr Leu Cys Ser Gly -10 Trp Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val Tyr Val Pro Tyr Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp Leu Val Ser Ile Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile Ala Ala Ile Thr Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp Glu Gly Ile Leu Gly Leu Ala Tyr Ala Glu Ile Ala Arg Pro Asp Asp Ser Pro Glu Pro Phe Phe Asp Ser Leu Val Lys Gln Thr His Val Pro Asn Leu Phe Ser 85 90 Leu Gln Leu Cys Gly Ala Gly Phe Pro Leu Asn Gln Ser Glu Val Leu 105 100 Ala Ser Val Gly Gly Ser Met Ile Ile Gly Gly Ile Asp His Ser Leu 120 Tyr Thr Gly Ser Leu Trp Tyr Thr Pro Ile Arg Arg Glu Trp Tyr Tyr 135 Glu Val Ile Ile Val Arg Val Glu Ile Asn Gly Gln Asp Leu Lys Met 150 155 Asp Cys Lys Glu Tyr Asn Tyr Asp Lys Ser Ile Val Asp Ser Gly Thr 170 165 Thr Asn Leu Arg Leu Pro Lys Lys Val Phe Glu Ala Ala Val Lys Ser 185 Ile Lys Ala Ala Ser Ser Thr Glu Lys Phe Pro Asp Gly Phe Trp Leu 200 Gly Glu Gln Leu Val Cys Trp Gln Ala Gly Thr Thr Pro Trp Asn Ile 215 220 Phe Pro Val Ile Ser Leu Tyr Leu Met Gly Glu Val Thr Asn Gln Ser 230 235 Phe Arg Ile Thr Ile Leu Pro Gln Gln Tyr Leu Arg Pro Val Glu Asp 250 Val Ala Thr Ser Gln Asp Asp Cys Tyr Lys Phe Ala Ile Ser Gln Ser 265 Ser Thr Gly Thr Val Met Gly Ala Val Ile Met Glu Gly Phe Tyr Val 280 Val Phe Asp Arg Ala Arg Lys Arg Ile Gly Phe Ala Val Ser Ala Cys 295 300 His Val His Asp Glu Phe Arg Thr Ala Ala Val Glu Gly Pro Phe Cys 310 315 His Leu Gly His Gly Arg Leu Trp Leu Gln His Ser Thr Asp Arg

330

<210> 168

<211> 138

<212> PRT

<213> Homo sapiens

325

<220>

<221> SIGNAL

<222> -47..-1

<400> 168 Met Glu Lys Phe Val Asp Pro Gly Asn His Asn Ser Gly Ile Asp Leu -40 -35 Leu Arg Thr Tyr Leu Trp Arg Cys Gln Phe Leu Leu Pro Phe Val Ser -25 -20 Leu Gly Leu Met Cys Phe Gly Ala Leu Ile Gly Leu Cys Ala Cys Ile -10 - 5. Cys Arg Ser Leu Tyr Pro Thr Ile Ala Thr Gly Ile Leu His Leu Leu Ala Gly Leu Cys Thr Leu Gly Ser Val Ser Cys Tyr Val Ala Gly Ile 25 Glu Leu Leu His Gln Lys Leu Glu Leu Pro Asp Asn Val Ser Gly Glu 40 Phe Gly Trp Ser Phe Cys Leu Ala Cys Val Ser Ala Pro Leu Gln Phe 55 60 Met Ala Ser Ala Leu Phe Ile Trp Ala Ala His Thr Asn Arg Arg Glu 75 Tyr Thr Leu Met Lys Ala Tyr Arg Val Ala

<210> 169 <211> 101 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -73..-1

<400> 169 Met Asn Leu Glu Arg Val Ser Asn Glu Glu Lys Leu Asn Leu Cys Arg

Lys Tyr Tyr Leu Gly Gly Phe Ala Phe Leu Pro Phe Leu Trp Leu Val -55 -50 -45Asn Ile Phe Trp Phe Tyr Arg Glu Ala Phe Leu Val Pro Ala Tyr Thr

-40 -35 -30
Glu Gln Ser Gln Ile Lys Gly Tyr Val Trp Arg Ser Ala Val Gly Phe
-25 -20 -15 -10

Leu Phe Trp Val Ile Val Leu Thr Ser Trp Ile Thr Ile Phe Gln Ile
-5 1 5

Tyr Arg Pro Arg Trp Gly Ala Leu Gly Asp Tyr Leu Ser Phe Thr Ile
10 20

Pro Leu Gly Thr Pro

<210> 170 <211> 252 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -68..-1

<400> 170
Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu
-65 -60 -55

Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val -45 Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser -30 Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro -15 -10 Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu 35 Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys 50 55 Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe 70 Arg Glu Asn Val Leu Arg Asn Leu Ala Asp Lys Ala Phe Asp Arg Pro 85 Ile Cys Glu Ala Leu Leu Asp Gln Arg Phe Phe Asn Gly Ile Gly Asn 100 Tyr Leu Arg Ala Glu Ile Leu Tyr Arg Leu Lys Ile Pro Pro Phe Glu 115 120 Lys Ala Arg Ser Val Leu Glu Ala Leu Gln Gln His Arg Pro Ser Pro 130 135 Glu Leu Thr Leu Ser Gln Lys Ile Arg Thr Lys Leu Gln Asn Ser Asp 150 145 Leu Leu Glu Leu Cys His Ser Val Pro Lys Glu Val Val Gln Leu Gly 165 Glu Ala Lys Asp Gly Ser Asn Leu Cys Phe Ser Lys 180

<210> 171 <211> 350 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -68..-1

<400> 171 Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu -60 Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val -45 Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser -30 Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro -10 Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His 20 Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe Arg Leu Lys Ile Pro Pro Phe Glu Lys Ala Arg Ser Val Leu Glu Ala

85 80 Leu Gln Gln His Arg Pro Ser Pro Glu Leu Thr Leu Ser Gln Lys Ile 100 105 Arg Thr Lys Leu Gln Asn Pro Asp Leu Leu Glu Leu Cys His Ser Val 115 Pro Lys Glu Val Asp Gln Leu Gly Gly Arg Gly Tyr Gly Ser Glu Ser 130 135 Gly Glu Glu Asp Phe Ala Ala Phe Arg Ala Trp Leu Arg Cys Tyr Gly 145 150 Met Pro Gly Met Ser Ser Leu Gln Asp Arg His Gly Arg Thr Ile Trp 165 Phe Gln Gly Asp Pro Gly Pro Leu Ala Pro Lys Gly Arg Lys Ser Arg 180 175 Lys Lys Lys Ser Lys Ala Thr Gln Leu Ser Pro Glu Asp Arg Val Glu 195 Asp Ala Leu Pro Pro Ser Lys Ala Pro Ser Lys Thr Arg Arg Ala Lys 215 Arg Asp Leu Pro Lys Arg Thr Ala Thr Gln Arg Pro Glu Gly Thr Ser 225 230 Leu Gln Gln Asp Pro Glu Ala Pro Thr Val Pro Lys Lys Gly Arg Arg 245 Lys Gly Arg Gln Ala Ala Ser Gly His Cys Arg Pro Arg Lys Val Lys 260 Ala Asp Ile Pro Ser Leu Glu Pro Glu Gly Thr Ser Ala Ser 275

<210> 172 <211> 390 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<222> -68..-1 <400> 172 Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu -60 Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val -45 Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser -30 -25 Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro -10 -15 Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His 20 Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu 35 Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys 50 Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe 70 65 Arg Glu Asn Val Leu Arg Asn Leu Ala Asp Lys Ala Phe Asp Arg Pro Ile Cys Glu Ala Leu Leu Asp Gln Arg Phe Phe Asn Gly Ile Gly Asn 100 Tyr Leu Arg Ala Glu Ile Leu Tyr Arg Leu Lys Ile Pro Pro Phe Glu

Lys Ala Arg Ser Val Leu Glu Ala Leu Gln Gln His Arg Pro Ser Pro 130 135 Glu Leu Thr Leu Ser Gln Lys Ile Arg Thr Lys Leu Gln Asn Pro Asp 145 150 Leu Leu Glu Leu Cys His Ser Val Pro Lys Glu Val Val Gln Leu Gly 165 Gly Arg Gly Tyr Gly Ser Glu Ser Gly Glu Glu Asp Phe Ala Ala Phe 180 Arg Ala Trp Leu Arg Cys Tyr Gly Met Pro Gly Met Ser Ser Leu Gln 195 Asp Arg His Gly Arg Thr Ile Trp Phe Gln Gly Asp Pro Gly Pro Leu 215 210 Ala Pro Lys Gly Arg Lys Ser Arg Lys Lys Lys Ser Lys Ala Thr Gln 225 230 Leu Ser Pro Glu Asp Arg Val Glu Asp Ala Leu Pro Pro Ser Lys Ala 245 Pro Ser Arg Thr Arg Arg Ala Lys Arg Asp Leu Pro Lys Arg Thr Ala 260 Thr Gln Arg Pro Glu Gly Thr Ser Leu Gln Gln Asp Pro Glu Ala Pro 275 280 Thr Val Pro Lys Lys Gly Arg Arg Lys Gly Arg Gln Ala Ala Ser Gly 290 295 His Cys Arg Pro Arg Lys Val Lys Ala Asp Ile Pro Ser Leu Glu Pro 305 Glu Gly Thr Ser Ala Ser 320

<210> 173 <211> 190 <212> PRT <213> Homo sapiens <220>

<400> 173

<221> SIGNAL <222> -82..-1

Met Tyr Val Trp Pro Cys Ala Val Leu Ala Gln Tyr Leu Trp Phe -75 His Arg Arg Ser Leu Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala Gly -60 Val Ser Leu Pro Gly Ile Leu Thr Ala Lys Cys Gly Ala Glu Val Ile -45 -40 Leu Ser Asp Ser Ser Glu Leu Pro His Cys Leu Glu Val Cys Arg Gln -25 Ser Cys Gln Met Asn Asn Leu Pro His Leu Gln Val Val Gly Leu Thr -10 Trp Gly His Ile Ser Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp Ile Ile Leu Ala Ser Asp Val Phe Phe Glu Pro Glu Asp Phe Glu Asp Ile 20 Leu Ala Thr Ile Tyr Phe Leu Met His Lys Asn Pro Lys Val Gln Leu Trp Ser Thr Tyr Gln Val Arg Ser Ala Asp Trp Ser Leu Glu Ala Leu Leu Tyr Lys Trp Asp Met Lys Cys Val His Ile Pro Leu Glu Ser Phe 70 Asp Ala Asp Lys Glu Asp Ile Ala Glu Ser Thr Leu Pro Gly Arg His 85

Thr Val Glu Met Leu Val Ile Ser Phe Ala Lys Asp Ser Leu

95

100

105

<210> 174 <211> 285 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -232..-1 <400> 174 Met Gly Cys Val Phe Gln Ser Thr Glu Asp Lys Arg Ile Phe Lys Ile -230 -225 -220 Asp Trp Thr Leu Ser Pro Gly Glu His Ala Lys Asp Glu Tyr Val Leu -215 -210 -205 Tyr Tyr Tyr Ser Asn Leu Ser Val Pro Ile Gly Arg Phe Gln Asn Arg -190 -185 -200 -195 Val His Leu Met Gly Asp Asn Leu Cys Asn Asp Gly Ser Leu Leu Leu -180 -175 Gln Asp Val Gln Glu Ala Asp Gln Gly Thr Tyr Ile Cys Glu Ile Arg -165 -160 Leu Lys Gly Glu Ser Gln Val Phe Lys Lys Ala Val Val Leu His Val -150 -145 -140 Leu Pro Glu Glu Pro Lys Glu Leu Met Val His Val Gly Gly Leu Ile -130 -125 Gln Met Gly Cys Val Phe Gln Ser Thr Glu Val Lys His Val Thr Lys -115 ~110 Val Glu Trp Ile Phe Ser Gly Arg Arg Ala Lys Glu Glu Ile Val Phe -100 -95 Arg Tyr Tyr His Lys Leu Arg Met Ser Ala Glu Tyr Ser Gln Ser Trp -85 -80 Gly His Phe Gln Asn Arg Val Asn Leu Val Gly Asp Ile Phe Arg Asn -65 Asp Gly Ser Ile Met Leu Gln Gly Val Arg Glu Ser Asp Gly Gly Asn -50 Tyr Thr Cys Ser Ile His Leu Gly Asn Leu Val Phe Lys Lys Thr Ile -35 -30 Val Leu His Val Ser Pro Glu Glu Pro Arg Thr Leu Val Thr Pro Ala -20 -15 Ala Leu Arg Pro Leu Val Leu Gly Gly Asn Gln Leu Val Ile Ile Val -5 1 Gly Ile Val Cys Ala Thr Ile Leu Leu Leu Pro Val Leu Ile Leu Ile 15 Val Lys Lys Thr Cys Gly Asn Lys Ser Ser Val Asn Ser Thr Val Leu 30 35 Val Lys Asn Thr Lys Lys Thr Asn Pro Lys Lys Lys

<210> 175

<211> 153

<212> PRT

<213> Homo sapiens

45

<400> 175

Met Gly Cys Val Phe Gln Ser Thr Val Asp Lys Cys Ile Phe Lys Ile 10 Asp Trp Thr Leu Ser Pro Gly Glu His Ala Lys Asp Glu Tyr Val Leu

25 Tyr Tyr Ser Asn Leu Ser Val Pro Ile Gly Arg Phe Gln Asn Arg 40 Val His Leu Met Gly Asp Ile Leu Cys Asn Asp Gly Ser Leu Leu Leu 55 Gln Asp Val Gln Glu Ala Asp Gln Gly Thr Tyr Ile Cys Glu Ile Arg 70 75 Leu Lys Gly Glu Ser Gln Val Phe Lys Lys Ala Val Val Leu His Val 90 Leu Pro Glu Glu Pro Lys Glu Leu Met Val His Val Gly Gly Leu Ile 105 Gln Met Gly Cys Val Phe Gln Ser Thr Glu Val Lys His Val Thr Lys 120 Val Glu Trp Ile Phe Ser Gly Arg Arg Ala Lys Val Thr Arg Arg Lys 135 His His Cys Val Arg Glu Gly Ser Gly 150

<210> 176 <211> 49 <212> PRT <213> Homo sapiens

<400> 176 Met Leu Xaa Gly Asp His Arg Ala Leu Leu Leu Lys Ile Trp Leu Leu 10 Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro Gly Arg Leu Val Val 25 Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro Ser Cys Pro Arg Phe Cys

<210> 177 <211> 99 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -24..-1

<400> 177

Met Lys Ser Ala Lys Leu Gly Phe Leu Leu Arg Phe Phe Ile Phe Cys -20 -15 Ser Leu Asn Thr Leu Leu Gly Gly Val Asn Lys Ile Ala Glu Lys Ile Cys Gly Asp Leu Lys Asp Pro Cys Lys Leu Asp Met Asn Phe Gly Ser Cys Tyr Glu Val His Phe Arg Tyr Phe Tyr Asn Arg Thr Ser Lys Arg Cys Glu Thr Phe Val Phe Ser Gly Cys Asn Gly Asn Leu Asn Asn 50 Phe Lys Leu Lys Ile Glu Arg Glu Val Ala Cys Val Ala Lys Tyr Lys

Pro Pro Arg

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<210> 178
<211> 95
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -37..-1
<400> 178
Met Ala Ser Pro Ala Val Asn Arg Trp Lys Arg Pro Arg Leu Lys Pro
                            -30
       -35
Val Trp Pro Arg Arg Leu Glu Ser Trp Leu Leu Leu Asp Ala Leu Leu
                                           -10
                       -15
Arg Leu Gly Asp Thr Lys Lys Lys Arg Gln Pro Glu Ala Ala Thr Lys
                   1
                                    5
Ser Cys Val Arg Ser Ser Cys Gly Gly Pro Ser Gly Asp Gly Pro Pro
                              20
Pro Cys Leu Gln Gln Pro Asp Pro Arg Ala Leu Ser Gln Ala Phe Ser
                           35
Arg Ser Phe Pro Leu Phe Pro Ser Leu Ala Gly Lys Ser Met Ile
                        50
<210> 179
<211> 121
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -23..-1
<400> 179
Met Met Leu Pro Gln Trp Leu Leu Leu Phe Leu Leu Phe Phe Phe
           -20
Leu Phe Leu Leu Thr Arg Gly Ser Leu Ser Pro Thr Lys Tyr Asn Leu
Leu Glu Leu Lys Glu Ser Cys Ile Arg Asn Gln Asp Cys Glu Thr Gly
                                        20
Cys Cys Gln Arg Ala Pro Asp Asn Cys Glu Ser His Cys Ala Glu Lys
Gly Ser Glu Gly Ser Leu Cys Gln Thr Gln Val Phe Phe Gly Gln Tyr
            45
                                50
Arg Ala Cys Pro Cys Leu Arg Asn Leu Thr Cys Ile Tyr Ser Lys Asn
                            65
Glu Lys Trp Leu Ser Ile Ala Tyr Gly Arg Cys Gln Lys Ile Gly Arg
                        80
                                           85
Gln Lys Leu Ala Lys Lys Met Phe Phe
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<210> 180 <211> 59 <212> PRT

<213> Homo sapiens

<400> 180

Met Ile Leu Cys Phe Leu Leu Pro His His Arg Leu Gln Glu Ala Arg

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<210> 181
<211> 86
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -14..-1
<400> 181
Met Val Ala Leu Asn Leu Ile Leu Val Pro Cys Cys Ala Ala Trp Cys
                -10
                                    -5
Asp Pro Arg Arg Ile His Ser Gln Asp Asp Val Pro Arg Ser Ser Ala
                           10
Ala Asp Thr Gly Ser Ala Met Gln Arg Arg Glu Ala Trp Ala Gly Trp
                       25
Arg Arg Ser Gln Pro Phe Ser Val Gly Leu Pro Ser Ala Glu Arg Leu
                  40
Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg Ser Leu Val Gly Glu Gly
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<210> 182 <211> 165 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -58..-1

Tyr Arg Ile Cys Asp Leu

<400> 182 Met Thr Arg Leu Cys Leu Pro Arg Pro Glu Ala Arg Glu Asp Pro Ile

 Pro
 Val
 Pro
 Pro
 Arg
 Gly
 Leu
 Gly
 Ala
 Gly
 Gly
 Ser
 Pro

 Val
 Arg
 Pro
 Val
 Ser
 Thr
 Trp
 Gly
 Pro
 Ser
 Trp
 Leu
 Leu

-50

 Leu Arg Ala Cys Gly Met Pro Leu Thr Leu Leu Gly Leu Ala Phe Cys 90
 95
 100

 Leu His Pro Trp Ala 105
 105
 105

<210> 183 <211> 80 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -35..-1 <400> 183 Met Pro Phe Gln Phe Gly Thr Gln Pro Arg Arg Phe Pro Val Glu Gly **-35 -30 -25** Gly Asp Ser Ser Ile Glu Leu Glu Pro Gly Leu Ser Ser Ala Ala -5 -10 -15 Cys Asn Gly Lys Glu Met Ser Pro Thr Arg Gln Leu Arg Arg Cys Pro Gly Ser His Cys Leu Thr Ile Thr Asp Val Pro Val Thr Val Tyr Ala 25 20 Thr Thr Arg Lys Pro Pro Ala Gln Ser Ser Lys Glu Met His Pro Lys

35

40

<210> 184 <211> 73 <212> PRT <213> Homo sapiens <220>

<221> SIGNAL <222> -21..-1

<210> 185
<211> 98
<212> PRT
<213> Homo sapiens
<400> 185
Met Leu Gly Ala Glu Thr Glu Glu Lys Leu Phe Asp Ala Pro Leu Ser

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<210> 186 <211> 112 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 186 Met Glu Ser Arg Val Leu Leu Arg Thr Phe Cys Leu Ile Phe Gly Leu Gly Ala Val Trp Gly Leu Gly Val Asp Pro Ser Leu Gln Ile Asp Val 1 Leu Thr Glu Leu Glu Leu Gly Glu Ser Thr Thr Gly Val Arg Gln Val 20 Pro Gly Leu His Asn Gly Thr Lys Ala Phe Leu Phe Gln Asp Thr Pro 35 Arg Ser Ile Lys Ala Ser Thr Ala Thr Ala Glu Gln Phe Phe Gln Lys Leu Arg Asn Lys His Glu Phe Thr Ile Leu Val Thr Leu Lys Gln Thr

His Leu Asn Ser Gly Val Ile Leu Ser Ile His His Leu Asp His Arg

<210> 187 <211> 70 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -44..-1 <400> 187 Met Cys Cys Tyr Cys Arg Ile Phe Cys Leu Arg Cys Thr Tyr Phe Pro -35 -40 Val His Cys Gly Met Cys Asn Leu Arg Tyr Phe Glu Phe Ser Thr Phe -20 -25 Leu Leu Ser Leu Ser Leu Ile Thr Tyr Cys Phe Trp Asp Pro Pro His -5 Arg Gly Ser His Ser Leu Ser Leu Glu His Thr Pro Leu Asp Phe Leu

15

65

10

Glu Trp Gly Leu Leu Arg

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<210> 188
<211> 92
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -13..-1
<400> 188
Met Leu Phe Ser Leu Ser Leu Leu Ser Asn Leu Asn Gln Ile Gly Ser
                                -5
            -10
Ser His Leu Asp Arg Pro His Ile Pro Gly Gln Ser Ala Gln Leu Phe
                        10
lle Tyr Gln Met Ser Ser Gln Gln Leu Gln Gln Gln Pro Ser Ala Asn
                                        30
                    25
Lys Lys Ala Gly Lys Ile His Asn Thr Pro Phe Ala Asn Gln Leu Asn
                40
Pro Thr Gln His Leu Ala Lys Pro Phe Gln Gln Ile Leu Pro Gly Arg
                                60
Gln Ser Gly Ser Leu Thr Ser Pro Phe Leu Ala Cys
        70
<210> 189
<211> 207
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -42..-1
<400> 189
Met His Ile Leu Gln Leu Leu Thr Thr Val Asp Asp Gly Ile Gln Ala
                             -35
        -40
Ile Val His Cys Pro Asp Thr Gly Lys Asp Ile Trp Asn Leu Leu Phe
                                             -15
                         -20
Asp Leu Val Cys His Glu Phe Cys Gln Ser Asp Asp Pro Pro Ile Ile
                     -5
                                         1
Leu Gln Glu Gln Lys Thr Val Leu Ala Ser Val Phe Ser Val Leu Ser
            10
                                 15
Ala Ile Tyr Ala Ser Gln Thr Glu Gln Glu Tyr Leu Lys Ile Glu Lys
                             30
        25
Val Asp Leu Pro Leu Ile Asp Ser Leu Ile Arg Val Leu Gln Asn Met
Glu Gln Cys Gln Lys Lys Pro Glu Asn Ser Ala Glu Ser Asn Thr Glu
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65

145

115

Glu Thr Lys Arg Thr Asp Leu Thr Gln Asp Asp Leu His Leu Lys Ile

Leu Lys Asp Ile Leu Cys Glu Phe Leu Ser Asn Ile Phe Gln Ala Leu 95 Thr Lys Glu Thr Val Ala Gln Gly Val Lys Glu Gly Gln Leu Ser Lys 110

Gln Lys Cys Ser Ser Ala Phe Gln Asn Leu Leu Pro Phe Tyr Ser Pro

Val Val Glu Asp Phe Ile Lys Ile Leu Arg Glu Val Asp Lys Ala Leu

Ala Asp Asp Leu Glu Lys Asn Phe Pro Ser Leu Lys Val Gln Thr

125

140

155 160 165

<210> 190 <211> 201 <212> PRT <213> Homo sapiens

<400> 190 Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala Leu Lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe Gln Glu Ile Pro Lys 25 Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln Leu Glu Lys Ile Glu 40 Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile Asn Ile Thr Glu Met Asn Lys Gln Ile Ser Leu Leu Thr Ser Ala Val Asn His Leu Lys Ala 70 Asn Val Lys Ser Ala Ala Asp Leu Ile Ser Leu Pro Thr Thr Val Glu 90 Gly Leu Gln Lys Ser Val Ala Ser Ile Gly Asn Thr Leu Asn Ser Val 105 100 His Leu Ala Val Glu Ala Leu Gln Lys Thr Val Asp Glu His Lys Lys 120 125 Thr Met Glu Leu Leu Gln Ser Asp Met Asn Gln His Phe Leu Lys Glu 135 140 Thr Pro Gly Ser Asn Gln Ile Ile Pro Ser Pro Ser Ala Thr Ser Glu 150 155 Leu Asp Asn Lys Thr His Ser Glu Asn Leu Lys Gln Met Gly Asp Arg 170 Ser Ala Thr Leu Lys Arg Gln Ser Leu Asp Gln Val Thr Asn Arg Thr

Ser Ala Thr Leu Lys Arg Gln Ser Leu Asp Gln Val Ti 180 185 Asp Thr Val Lys Ile Gln Lys Lys Lys 195 200

<210> 191 <211> 379 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -37..-1

Ala Leu Leu Leu Ser Ile Tyr Phe Tyr Tyr Ser Leu Pro Asn Ala

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70 60 65 Val Gly Pro Pro Phe Thr Trp Met Leu Ala Leu Leu Gly Leu Ser Gln Ala Leu Asn Ile Leu Leu Gly Leu Lys Gly Leu Ala Pro Ala Glu Ile 100 Ser Ala Val Cys Glu Lys Gly Asn Phe Asn Val Ala His Gly Leu Ala 115 Trp Ser Tyr Tyr Ile Gly Tyr Leu Arg Leu Ile Leu Pro Glu Leu Gln 130 135 Ala Arg Ile Arg Thr Tyr Asn Gln His Tyr Asn Asn Leu Leu Arg Gly 150 145 Ala Val Ser Gln Arg Leu Tyr Ile Leu Leu Pro Leu Asp Cys Gly Val 165 Pro Asp Asn Leu Ser Met Ala Asp Pro Asn Ile Arg Phe Leu Asp Lys 180 Leu Pro Gln Gln Thr Gly Asp Arg Ala Gly Ile Lys Asp Arg Val Tyr 195 Ser Asn Ser Ile Tyr Glu Leu Leu Glu Asn Gly Gln Arg Ala Gly Thr 210 215 Cys Val Leu Glu Tyr Ala Thr Pro Leu Gln Thr Leu Phe Ala Met Ser 225 230 Gln Tyr Ser Gln Ala Gly Phe Ser Arg Glu Asp Arg Leu Glu Gln Ala 240 245 Lys Leu Phe Cys Arg Thr Leu Glu Asp Ile Leu Ala Asp Ala Pro Glu 260 Ser Gln Asn Asn Cys Arg Leu Ile Ala Tyr Gln Glu Pro Ala Asp Asp 275 Ser Ser Phe Ser Leu Ser Gln Glu Val Leu Arg His Leu Arg Gln Glu 290 295 Glu Lys Glu Glu Val Thr Val Gly Ser Leu Lys Thr Ser Ala Val Pro 305 310 Ser Thr Ser Thr Met Ser Gln Glu Pro Glu Leu Leu Ser Gly Met 320 325 Gly Lys Pro Leu Pro Leu Arg Thr Asp Phe Ser 335

<210> 192 <211> 112 <212> PRT <213> Homo sapiens

<400> 192 Met Pro Ser Glu Gly Arg Cys Trp Glu Thr Leu Lys Ala Leu Arg Ser 10 Ser Asp Lys Gly Arg Leu Cys Tyr Tyr Arg Asp Trp Leu Leu Arg Arg 25 Glu Asp Val Leu Glu Glu Cys Met Ser Leu Pro Lys Leu Ser Ser Tyr 40 Ser Gly Trp Val Val Glu His Val Leu Pro His Met Gln Glu Asn Gln 55 Pro Leu Ser Glu Thr Ser Pro Ser Ser Thr Ser Ala Ser Ala Leu Asp 70 75 Gln Pro Ser Phe Val Pro Lys Ser Pro Asp Ala Ser Ser Ala Phe Ser 90 85 Pro Ala Ser Pro Ala Thr Pro Asn Gly Thr Lys Gly Lys Lys Lys 105

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<211> 43
<212> PRT
<213> Homo sapiens
<400> 193
Ser Leu Pro Gln Ala Leu Trp Phe Gln Phe Phe Tyr His Ser Gly Ser
                                                        15
Ser Leu Glu Ser Pro Gly Met Leu Asn Gly Pro Phe Gln His Arg Asn
Ser Arg Ile Met Thr His Arg Ser Ala Glu Lys
       35
<210> 194
<211> 51
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -16..-1
<400> 194
Met Leu Arg Ile Ala Leu Thr Leu Ile Pro Ser Met Leu Ser Arg Ala
                        -10
    -15
Ala Gly Trp Cys Trp Tyr Lys Glu Pro Thr Gln Gln Phe Ser Tyr Leu
Cys Leu Pro Cys Leu Ser Trp Asn Lys Lys Gly Asn Val Leu Gln Leu
                               25
Pro Asn Phe
        35
<210> 195
<211> 244
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -18..-1
<400> 195
Met Ala Asn Pro Lys Leu Leu Gly Leu Glu Leu Ser Glu Ala Glu Ala
            -15
Ile Gly Ala Asp Ser Ala Arg Phe Glu Glu Leu Leu Gln Ala Ser
Lys Glu Leu Gln Gln Ala Gln Thr Thr Arg Pro Glu Ser Thr Gln Ile
                    20
                                        25
Gln Pro Gln Pro Gly Phe Cys Ile Lys Thr Asn Ser Ser Glu Gly Lys
Val Phe Ile Asn Ile Cys His Ser Pro Ser Ile Pro Pro Pro Ala Asp
                                 55
Val Thr Glu Glu Glu Leu Leu Gln Met Leu Glu Glu Asp Gln Ala Gly
                             70
Phe Arg Ile Pro Met Ser Leu Gly Glu Pro His Ala Glu Leu Asp Ala
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Lys Gly Gln Gly Cys Thr Ala Tyr Asp Val Ala Val Asn Ser Asp Phe

Tyr Arg Arg Met Gln Asn Ser Asp Phe Leu Arg Glu Leu Val Ile Thr

105

115 120 Ile Ala Arg Glu Gly Leu Glu Asp Ile Tyr Asn Leu Gln Leu Asn Pro 135 140 130 Glu Trp Arg Met Met Lys Asn Arg Pro Phe Met Gly Ser Ile Ser Gln 150 Gln Asn Ile Arg Ser Glu Gln Arg Pro Arg Ile Gln Glu Leu Gly Asp 170 165 Leu Tyr Thr Pro Ala Pro Gly Arg Ala Glu Ser Gly Pro Glu Lys Pro 180 185 His Leu Asn Leu Trp Leu Glu Ala Pro Asp Leu Leu Ala Glu Val 195 200 Asp Leu Pro Lys Leu Asp Gly Ala Leu Gly Leu Ser Leu Glu Ile Gly 210 215 Arg Thr Ala Trp 225

<210> 196 <211> 353 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -34..-1

<400> 196

Met Glu Arg Gly Leu Lys Ser Ala Asp Pro Arg Asp Gly Thr Gly Tyr -30 -25 Thr Gly Trp Ala Gly Ile Ala Val Leu Tyr Leu His Leu Tyr Asp Val -10 Phe Gly Asp Pro Ala Tyr Leu Gln Leu Ala His Gly Tyr Val Lys Gln 10 Ser Leu Asn Cys Leu Thr Lys Arg Ser Ile Thr Phe Leu Cys Gly Asp 20 25 Ala Gly Pro Leu Ala Val Ala Ala Val Leu Tyr His Lys Met Asn Asn 40 Glu Lys Gln Ala Glu Asp Cys Ile Thr Arg Leu Ile His Leu Asn Lys 55 Ile Asp Pro His Ala Pro Asn Glu Met Leu Tyr Gly Arg Ile Gly Tyr 70 Ile Tyr Ala Leu Leu Phe Val Asn Lys Asn Phe Gly Val Glu Lys Thr 85 90 Pro Gln Ser His Ile Gln Gln Ile Cys Glu Thr Ile Leu Thr Ser Gly 100 105 Glu Asn Leu Ala Arg Lys Arg Asn Phe Thr Ala Lys Ser Pro Leu Met 120 115 Tyr Glu Trp Tyr Gln Glu Tyr Tyr Val Gly Ala Ala His Gly Leu Ala 130 135 140 Gly Ile Tyr Tyr Leu Met Gln Pro Ser Leu Gln Val Ser Gln Gly 150 155 Lys Leu His Ser Leu Val Lys Pro Ser Val Asp Tyr Val Cys Gln Leu 165 170 Lys Phe Pro Ser Gly Asn Tyr Pro Pro Cys Ile Gly Asp Asn Arg Asp 185 180 Leu Leu Val His Trp Cys His Gly Ala Pro Gly Val Ile Tyr Met Leu 200 Ile Gln Ala Tyr Lys Val Phe Arg Glu Glu Lys Tyr Leu Cys Asp Ala 215 Tyr Gln Cys Ala Asp Val Ile Trp Gln Tyr Gly Leu Leu Lys Lys Gly 230

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```
Tyr Gly Leu Cys His Gly Ser Ala Gly Asn Ala Tyr Ala Phe Leu Thr
240

Leu Tyr Asn Leu Thr Gln Asp Met Lys Tyr Leu Tyr Arg Ala Cys Lys
255

Phe Ala Glu Trp Cys Leu Glu Tyr Gly Glu His Gly Cys Arg Thr Pro
275

Asp Thr Pro Phe Ser Leu Phe Glu Gly Met Ala Gly Thr Ile Tyr Phe
290

Leu Ala Asp Leu Leu Val Pro Thr Lys Ala Arg Phe Pro Ala Phe Glu
305

Leu Ala Asp Leu Leu Slu Tyr Thr Lys Ala Arg Phe Pro Ala Phe Glu
305

Leu Ala Asp Leu Leu Val Pro Thr Lys Ala Arg Phe Pro Ala Phe Glu
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<210> 197 <211> 30 <212> PRT <213> Homo sapiens

<210> 198
<211> 112
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -48..-1

<400> 198

<210> 199
<211> 54
<212> PRT
<213> Homo sapiens
<400> 199
Glu Ile Ala Gly Tyr Gly Ala Glu Gly Phe Ser Ser Val Leu Gly Tyr

 Pro Arg Trp His Arg Leu Pro Pro Gln Ser Leu Gln His His Gln Tyr

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 25
 30

 Cys Gln Arg Arg Trp Pro Asp Arg Arg Cys Leu Gln Ser His Thr Gln
 35
 40
 45

 Ser Ser Gly His Leu Pro
 50
 45
 45

<210> 200 <211> 151 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 200 Met Ala Ala Ser Thr Ser Met Xaa Pro Val Ala Val Thr Ala Ala Val -15 -10 Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu Arg Glu Ile 1 Lys Lys Gln Leu Leu Leu Ile Ala Gly Leu Thr Arg Glu Arg Gly Leu 20 Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser Leu Pro Ala 40 35 Leu Pro Xaa Gly Gln Leu Gln Pro Pro Pro Pro Ile Thr Glu Glu Asp . ..55 50 Ala Gln Asp Met Asp Ala Tyr Thr Leu Ala Lys Ala Tyr Phe Asp Val 70 65 Lys Glu Tyr Asp Arg Ala Ala His Phe Leu His Gly Cys Asn Ser Lys 85 80 Lys Ala Tyr Phe Leu Tyr Met Tyr Ser Arg Tyr Leu Val Arg Ala Ile 100 Leu Lys Cys His Ser Ala Phe Ser Glu Thr Ser Ile Phe Arg Thr Asn 110 Gly Lys Val Lys Ser Phe Lys

<210> 201 <211> 228 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -25..-1 <400> 201 Met Ser Met Ala Val -25

125

 Met
 Ser
 Met
 Ala
 Val
 Glu
 Thr
 Phe
 Gly
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 Met
 Ala
 Thr
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 Gly
 Phe
 Gly
 Phe
 Met
 Ala
 Thr
 Val
 Thr
 Leu
 Pro
 Asn
 Ser
 Tyr
 Trp
 Arg
 Val
 Ser

 Thr
 Val
 His
 Gly
 Asn
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 Thr

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```
Leu Met Ile Thr Ala Ile Leu Leu Gly Phe Leu Gly Leu Leu Gly
Ile Ala Gly Leu Arg Cys Thr Asn Ile Gly Gly Leu Glu Leu Ser Arg
                              80
Lys Ala Lys Leu Ala Ala Thr Ala Gly Ala Pro His Ile Leu Ala Gly
                          95
Ile Cys Gly Met Val Ala Ile Ser Trp Tyr Ala Phe Asn Ile Thr Arg
                      110
Asp Phe Phe Asp Pro Leu Tyr Pro Gly Thr Lys Tyr Glu Leu Gly Pro
                  125
                                  130
Ala Leu Tyr Leu Gly Trp Ser Ala Ser Leu Ile Ser Ile Leu Gly Gly
                        145
Leu Cys Leu Cys Ser Ala Cys Cys Cys Gly Ser Asp Glu Asp Pro Ala
                             160
Ala Ser Ala Arg Arg Pro Tyr Gln Ala Pro Val Ser Val Met Pro Val
                          175
Ala Thr Ser Asp Gln Glu Gly Asp Ser Ser Phe Gly Lys Tyr Gly Arg
                      190
Asn Ala Tyr Val
200
```

<210> 202 <211> 64 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -47..-1

Met His Gly Phe Glu Ile Ile Ser Leu Lys Glu Glu Ser Pro Leu Gly
-45

Lys Val Ser Gln Gly Pro Leu Phe Asn Val Thr Ser Gly Ser Ser Ser
-30

Pro Val Thr Trp Leu Gly Leu Leu Ser Phe Gln Asn Leu His Cys Phe
-15

Pro Asp Leu Pro Thr Glu Met Pro Leu Arg Ala Lys Gly Val Asn Thr

<210> 203
<211> 146
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -31..-1
<400> 203
Met Met Trp Gln Lys

Met Met Trp Gln Lys Tyr Ala Gly Ser Arg Arg Ser Met Pro Leu Gly
-30
-25
-20

Ala Arg Ile Leu Phe His Gly Val Phe Tyr Ala Gly Gly Phe Ala Ile
-15
-10
-5
1

Val Tyr Tyr Leu Ile Gln Lys Phe His Ser Arg Ala Leu Tyr Tyr Lys
5
10
Leu Ala Val Glu Gln Leu Gln Ser His Pro Glu Ala Gln Glu Ala Leu
20
25
30

<210> 204 <211> 87 <212> PRT <213> Homo sapiens

<400> 204

 Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly Ser Leu

 1
 5
 10
 15

 Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His Leu
 20
 25
 30

 Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro Glu
 35
 40
 45

 Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln Ser
 50
 60

 Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu Leu
 65
 70
 75
 80

 Glu Val Asp Asp Trp Glu Phe
 85
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<210> 205
<211> 40
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -27...-1

Met Gly Ser Leu Ser Gly Leu Arg Leu Ala Ala Gly Ser Cys Phe Arg

<210> 206 <211> 154 <212> PRT <213> Homo sapiens

<400> 206

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10
Leu Cys Glu Arg Asp Val Ser Ser Ser Leu Arg Leu Thr Arg Ser Ser
                               25
Asp Leu Lys Arg Ile Asn Gly Phe Cys Thr Lys Pro Gln Glu Ser Pro
                            40
Gly Ala Pro Ser Arg Thr Tyr Asn Arg Val Pro Leu His Lys Pro Thr
Asp Trp Gln Lys Lys Ile Leu Ile Trp Ser Gly Arg Phe Lys Lys Glu
Asp Glu Ile Pro Glu Thr Val Ser Leu Glu Met Leu Asp Ala Ala Lys
                                    90
Asn Lys Met Arg Val Lys Ser Ser Tyr Leu Met Ile Ala Leu Thr Val
           100
                               105
Val Gly Cys Ile Phe Met Val Ile Glu Gly Lys Lys Ala Ala Gln Arg
                            120
His Glu Thr Leu Thr Ser Leu Asn Leu Glu Lys Lys Ala Arg Leu Lys
                       135
Glu Glu Ala Ala Met Lys Ala Lys Thr Glu
```

<210> 207 <211> 101 <212> PRT <213> Homo sapiens

<400> 207

 Met
 Val
 Cys
 Glu
 Lys
 Cys
 Glu
 Lys
 Lys
 Lys
 Leu
 Gly
 Thr
 Val
 Ile
 Thr
 Pro

 Asp
 Thr
 Thr
 Thr
 Thr
 Thr
 Glu
 Ser
 Gly
 Arg
 Arg

 Lys
 Leu
 Asp
 Lys
 Ala
 Leu
 Thr
 Ser
 Lys
 Ala
 Arg
 Phe
 Arg
 Arg
 Ile
 Cys
 Lys
 Ser
 Ser
 Ser
 Ser
 Thr
 Cys
 Arg
 Ile
 Cys
 Lys
 Ser
 Ser

<210> 208 <211> 456 <212> PRT <213> Homo sapiens <220> <221> SIGNAL

<400> 208

<222> -22..-1

Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val Ala Ala Gly
-20
-15
-10

Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser Ser Gln Asn
-5
1
Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg Ala Leu Glu
15
20
25

Ala Ala Ser Leu Ser Gln His Pro Pro Ser Leu Cys Ile Ser Asp Ser

```
30
                              35
Glu Glu Glu Glu Glu Arg Lys Lys Cys Pro Lys Lys Ala Ser
                         50
Phe Ala Ser Ala Ser Ala Glu Val Gly Lys Lys Gly Lys Lys Cys
Gln Lys Gln Gly Pro Pro Cys Ser Asp Ser Glu Glu Glu Val Glu Arg
Lys Lys Lys Cys His Lys Gln Ala Leu Val Gly Ser Asp Ser Ala Glu
                                  100
Asp Glu Lys Arg Lys Arg Lys Cys Gln Lys His Ala Pro Ile Asn Ser
           110
                              115
Ala Gln His Leu Asp Asn Val Asp Gln Thr Gly Pro Lys Ala Trp Lys
                          130
Gly Ser Thr Thr Asn Asp Pro Pro Lys Gln Ser Pro Gly Ser Thr Ser
                      145
                                          150
Pro Lys Pro Pro His Thr Leu Ser Arg Lys Gln Trp Arg Asn Arg Gln
                  160
                                     165
Lys Asn Lys Arg Arg Cys Lys Asn Lys Phe Gln Pro Pro Gln Val Pro
              175
                                 180
Asp Gln Ala Pro Ala Glu Ala Pro Thr Glu Lys Thr Glu Val Ser Pro
                              195
Val Pro Arg Thr Asp Ser His Gly Ala Arg Ala Gly Ala Leu Arg Ala
                          210
Arg Met Ala Gln Arg Leu Asp Gly Ala Arg Phe Arg Tyr Leu Asn Glu
                      225
                                          230
Gln Leu Tyr Ser Gly Pro Ser Ser Ala Ala Gln Arg Leu Phe Gln Glu
                   240
                                      245
Asp Pro Glu Ala Phe Leu Leu Tyr His Arg Gly Phe Gln Ser Gln Val
               255
                                  260
Lys Lys Trp Pro Leu Gln Pro Val Asp Arg Ile Ala Arg Asp Leu Arg
           270
                              275
Gln Arg Pro Ala Ser Leu Val Val Ala Asp Phe Gly Cys Gly Asp Cys
                           290
Arg Leu Ala Ser Ser Ile Arg Asn Pro Val His Cys Phe Asp Leu Ala
                       305
                                          310
Ser Leu Asp Pro Arg Val Thr Val Cys Asp Met Ala Gln Val Pro Leu
                   320
                                      325
Glu Asp Glu Ser Val Asp Val Ala Val Phe Cys Leu Ser Leu Met Gly
              335
                                  340
Thr Asn Ile Arg Asp Phe Leu Glu Glu Ala Asn Arg Val Leu Lys Pro
           350
                              355
Gly Gly Leu Leu Lys Val Ala Glu Val Ser Ser Arg Phe Glu Asp Val
                          370
Arg Thr Phe Leu Arg Ala Val Thr Lys Leu Gly Phe Lys Ile Val Ser
                      385
Lys Asp Leu Thr Asn Ser His Phe Phe Leu Phe Asp Phe Gln Lys Thr
                  400
                                     405
Gly Pro Pro Leu Val Gly Pro Lys Ala Gln Leu Ser Gly Leu Gln Leu
               415
                                  420
Gln Pro Cys Leu Tyr Lys Arg Arg
           430
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<210> 209
<211> 98

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -17..-1

PCT/IB98/02122

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      <400> 209

      Met Pro Ser Ser Phe Phe Leu Leu Leu Gln Phe Phe Leu Arg Ile Asp -15

      Gly Val Leu Ile Arg Met Asn Asp Thr Arg Leu Tyr His Glu Ala Asp 1

      Lys Thr Tyr Met Leu Arg Glu Tyr Thr Ser Arg Glu Ser Lys Ile Ser 20

      Ser Leu Met His Val Pro Pro Ser Leu Phe Thr Glu Pro Asn Glu Ile 35

      Ser Gln Tyr Leu Pro Ile Lys Glu Ala Val Cys Glu Lys Leu Ile Phe 50

      Pro Glu Arg Ile Asp Pro Asn Pro Ala Asp Ser Gln Lys Ser Thr Gln 65

      Val Glu 80
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<210> 210 <211> 83 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -29..-1

<400> 210

Asn Ala Ser

 Met
 Thr
 Leu
 Leu
 Ser
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 Ala
 Ala
 Phe
 Thr
 Ala
 Ala
 Phe
 Ser
 Val
 Leu
 Leu
 Phe
 Classification
 Ala
 Ala
 Ala
 Phe
 Ser
 Ala
 Ala
 Leu
 Ala
 Ser
 Val
 Phe
 Phe
 Ala
 Ala
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 Phe

<210> 211 <211> 229 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -23..-1

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            45
Gly Lys Thr Leu Val Phe Glu Gln Arg Lys Ser Asp Gly Val His Thr
                           65
Val Glu Thr Glu Val Gly Asp Tyr Met Phe Cys Phe Asp Asn Thr Phe
Ser Thr Ile Ser Glu Lys Val Ile Phe Phe Glu Leu Ile Leu Asp Asn
                                       100
Met Gly Glu Gln Ala Gln Glu Gln Glu Asp Trp Lys Lys Tyr Ile Thr
               110
                                   115
Gly Thr Asp Ile Leu Asp Met Lys Leu Glu Asp Ile Leu Glu Ser Ile
                               130
Ser Ser Ile Lys Ser Arg Leu Ser Lys Ser Gly His Ile Gln Ile Leu
                           145
                                               150
Leu Arg Ala Phe Glu Ala Arg Asp Arg Asn Ile Gln Glu Ser Asn Phe
                      160
                                           165
Asp Arg Val Asn Phe Trp Ser Met Val Asn Leu Val Val Met Val Val
                  175
                                      180
Val Ser Ala Ile Gln Val Tyr Met Leu Lys Ser Leu Phe Glu Asp Lys
               190
                                  195
Arg Lys Ser Arg Thr
            205
```

<210> 212 <211> 152 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 212 Met Ala Gln Leu Gly Ala Val Val Ala Val Ala Ser Ser Phe Phe Cys Ala Ser Leu Phe Ser Ala Val His Lys Ile Glu Glu Gly His Ile Gly Val Tyr Tyr Arg Gly Gly Ala Leu Leu Thr Ser Thr Ser Gly Pro Gly 20 Phe His Leu Met Leu Pro Phe Ile Thr Ser Tyr Lys Ser Val Gln Thr 35 Thr Leu Gln Thr Asp Glu Val Lys Asn Val Pro Cys Gly Thr Ser Gly 50 Gly Val Met Ile Tyr Phe Asp Arg Ile Glu Val Val Asn Phe Leu Val Pro Asn Ala Val His Asp Ile Val Lys Asn Tyr Thr Ala Asp Tyr Asp Lys Ala Leu Ile Phe Asn Lys Ile His His Glu Leu Asn Gln Phe Cys 100 Ser Val His Thr Leu Gln Glu Val Tyr Ile Glu Leu Phe Gly Leu Glu 115 Asn Asp Phe Ser Gln Glu Ser Ser 125

<210> 213

<211> 179

<212> PRT

<213> Homo sapiens

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<220>
<221> SIGNAL
<222> -54..-1
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<400> 213 Met Ala Ala Ser Glu Ala Ala Val Val Ser Ser Pro Ser Leu Lys Thr -50 -45 Asp Thr Ser Pro Val Leu Glu Thr Ala Gly Thr Val Ala Ala Met Ala Ala Thr Pro Ser Ala Arg Ala Ala Ala Ala Val Val Ala Ala Ala -15 Arg Thr Gly Ser Glu Ala Arg Val Ser Lys Ala Ala Leu Ala Thr Lys Leu Leu Ser Leu Ser Gly Val Phe Ala Val His Lys Pro Lys Gly Pro 20 Thr Ser Ala Glu Leu Leu Asn Arg Leu Lys Glu Lys Leu Leu Ala Glu 35 Ala Gly Met Pro Ser Pro Glu Trp Thr Lys Arg Lys Lys Gln Thr Leu 50 Lys Ile Gly His Gly Gly Thr Leu Asp Ser Ala Ala Arg Gly Val Leu Val Val Gly Ile Gly Ser Gly Thr Lys Met Leu Thr Ser Met Leu Ser 85 80 Gly Ser Lys Arg Tyr Thr Ala Ile Gly Glu Leu Gly Lys Ala Thr Asp 100 95 Thr Leu Asp Ser Thr Gly Lys Val Thr Glu Glu Lys Pro Tyr Gly Met 115

<210> 214 <211> 269 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -92..-1

125

Asn Leu Ile

<400> 214 Met Ile Thr His Val Thr Leu Glu Asp Ala Leu Ser Asn Val Asp Leu -85 Leu Glu Glu Leu Pro Leu Pro Asp Gln Gln Pro Cys Ile Glu Pro Pro -70 Pro Ser Ser Ile Met Tyr Gln Ala Asn Phe Asp Thr Asn Phe Glu Asp -50 -55 Arg Asn Ala Phe Val Thr Gly Ile Ala Arg Tyr Ile Glu Gln Ala Thr -35 -40 Val His Ser Ser Met Asn Glu Met Leu Glu Glu Gly His Glu Tyr Ala -20 Val Met Leu Tyr Thr Trp Arg Ser Cys Ser Arg Ala Ile Pro Gln Val -5 Lys Cys Asn Glu Gln Pro Asn Arg Val Glu Ile Tyr Glu Lys Thr Val 15 Glu Val Leu Glu Pro Glu Val Thr Lys Leu Met Lys Phe Met Tyr Phe 30 Gln Arg Lys Ala Ile Glu Arg Phe Cys Ser Glu Val Lys Arg Leu Cys 45 His Ala Glu Arg Arg Lys Asp Phe Val Ser Glu Ala Tyr Leu Leu Thr 60

Leu Gly Lys Phe Ile Asn Met Phe Ala Val Leu Asp Glu Leu Lys Asn 75 Met Lys Cys Ser Val Lys Asn Asp His Ser Ala Tyr Lys Arg Ala Ala 95 90 Gln Phe Leu Arg Lys Met Ala Asp Pro Gln Ser Ile Gln Glu Ser Gln 105 110 Asn Leu Ser Met Phe Leu Ala Asn His Asn Arg Ile Thr Gln Cys Leu 125 120 His Gln Gln Leu Glu Val Ile Pro Gly Tyr Glu Glu Leu Leu Ala Asp 140 145 Ile Val Asn Ile Cys Val Asp Tyr Tyr Glu Asn Lys Met Tyr Leu Thr 155 Pro Ser Glu Lys His Met Leu Leu Lys Val Lys Leu Pro 170

<210> 215 <211> 135 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -22..-1

<400> 215

 Met
 Gln
 Thr
 Val
 Tyr
 Tyr
 Gly
 Ser
 Leu
 Gly
 Leu
 Trp
 Leu
 Ala
 Leu
 Val

 Asp
 Gly
 Leu
 Val
 Arg
 Ser
 Ser
 Pro
 Ser
 Leu
 Asp
 Gln
 Met
 Phe
 Asp
 Ala

 Glu
 Ile
 Leu
 Gly
 Phe
 Ser
 Thr
 Pro
 Pro
 Gly
 Arg
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 Ser
 Met
 Asp

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 Ile
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 Gly
 Phe
 Ser
 Thr
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 Ala
 Leu
 Gly
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 Tyr
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<210> 216 <211> 67 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -38..-1

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<210> 217 <211> 125 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -54..-1

<400> 217

Met Ala Asp Glu Glu Leu Glu Ala Leu Arg Arg Gln Arg Leu Ala Glu -45 -50 Leu Gln Ala Lys His Gly Asp Pro Gly Asp Ala Ala Gln Gln Glu Ala -30 -25 -35 Lys His Arg Glu Ala Glu Met Arg Asn Ser Ile Leu Ala Gln Val Leu -20 -15 -10 Asp Gln Ser Ala Arg Ala Arg Leu Ser Asn Leu Ala Leu Val Lys Pro 5 Glu Lys Thr Lys Ala Val Glu Asn Tyr Leu Ile Gln Met Ala Arg Tyr 15 Gly Gln Leu Ser Glu Lys Val Ser Glu Gln Gly Leu Ile Glu Ile Leu Lys Lys Val Ser Gln Gln Thr Glu Lys Thr Thr Thr Val Lys Phe Asn Arg Arg Lys Val Met Asp Ser Asp Glu Asp Asp Asp Tyr

<210> 218 <211> 376 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -21..-1 <400> 218 Met Gly His Arg Phe Leu Arg Gly Leu Leu Thr Leu Leu Leu Pro Pro -15 Pro Pro Leu Tyr Thr Arg His Arg Met Leu Gly Pro Glu Ser Val Pro Pro Pro Lys Arg Ser Arg Ser Lys Leu Met Ala Pro Pro Arg Ile Gly 20 Thr His Asn Gly Thr Phe His Cys Asp Glu Ala Leu Ala Cys Ala Leu 35 Leu Arg Leu Leu Pro Glu Tyr Arg Asp Ala Glu Ile Val Arg Thr Arg 50 Asp Pro Glu Lys Leu Ala Ser Cys Asp Ile Val Val Asp Val Gly Gly 70 Glu Tyr Asp Pro Arg Arg His Arg Tyr Asp His His Gln Arg Ser Phe

Thr Glu Thr Met Ser Ser Leu Ser Pro Gly Arg Pro Trp Gln Thr Lys

100 Leu Ser Ser Ala Gly Leu Ile Tyr Leu His Phe Gly His Lys Leu Leu 115 120 Ala Gln Leu Leu Gly Thr Ser Glu Glu Asp Ser Met Val Gly Thr Leu 130 135 Tyr Asp Lys Met Tyr Glu Asn Phe Val Glu Glu Val Asp Ala Val Asp 145 150 Asn Gly Ile Ser Gln Trp Ala Glu Gly Glu Pro Arg Tyr Ala Leu Thr 160 165 Thr Thr Leu Ser Ala Arg Val Ala Arg Leu Asn Pro Thr Trp Asn His 175 180 Pro Asp Gln Asp Thr Glu Ala Gly Phe Lys Arg Ala Met Asp Leu Val 195 200 Gln Glu Glu Phe Leu Gln Arg Leu Asp Phe Tyr Gln His Ser Trp Leu 215 210 Pro Ala Arg Ala Leu Val Glu Glu Ala Leu Ala Gln Arg Phe Gln Val 230 Asp Pro Ser Gly Glu Ile Val Glu Leu Ala Lys Gly Ala Cys Pro Trp Lys Glu His Leu Tyr His Leu Glu Ser Gly Leu Ser Pro Pro Val Ala 260 Ile Phe Phe Val Ile Tyr Thr Asp Gln Ala Gly Gln Trp Arg Ile Gln 275 Cys Val Pro Lys Glu Pro His Ser Phe Gln Ser Arg Leu Pro Leu Pro 290 295 Glu Pro Trp Arg Gly Leu Arg Asp Glu Ala Leu Asp Gln Val Ser Gly 305 310 Ile Pro Gly Cys Ile Phe Val His Ala Ser Gly Phe Ile Gly Gly His 325 320 Arg Thr Arg Glu Gly Ala Leu Ser Met Ala Arg Ala Thr Leu Ala Gln 335 340 Arg Ser Tyr Leu Pro Gln Ile Ser 350

<210> 219 <211> 211

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -30..-1

<400> 219

 Met
 Gly
 Glu
 Ala
 Ser
 Pro
 Pro
 Ala
 Pro
 Ala
 Arg
 Arg
 His
 Leu
 Leu
 Val

 -30
 -25
 -25
 -25
 -20
 -20
 -15

 Leu
 Leu
 Leu
 Ser
 Thr
 Leu
 Val
 Ile
 Pro
 Ser
 Ala
 Ala
 Ala
 Pro
 Pro
 Ser
 Ala
 Ala
 Ala
 Pro
 Ser
 Ser
 Ala
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 Leu
 Fro
 Ser
 Leu
 Gly
 Leu
 Fro
 Ser
 Leu
 Pro
 Leu
 Pro
 Dro
 Dro
 Incompatible
 Incompatible</

<210> 220

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<211> 154
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -60..-1
<400> 220
Met Gly Ser Lys Cys Lys Gly Gly Pro Asp Glu Asp Ala Val Glu
                   -55
                                       -50
Arg Gln Arg Arg Gln Lys Leu Leu Leu Ala Gln Leu His His Arg Lys
               -40
                                   -35 .. -30
Arg Val Lys Ala Ala Gly Gln Ile Gln Ala Trp Trp Arg Gly Val Leu
                               -20
Val Arg Arg Thr Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln
Cys Trp Trp Arg Thr Leu Val Gln Arg Arg Ile Arg Gln Arg Arg Gln
                  10
                                      15
Ala Leu Leu Arg Val Tyr Val Ile Gln Glu Gln Ala Thr Val Lys Leu
               25
                                  30
Gln Ser Cys Ile Arg Met Trp Gln Cys Arg Gln Cys Tyr Arg Gln Met
                              45
Cys Asn Ala Leu Cys Leu Phe Gln Val Pro Glu Ser Ser Leu Ala Phe
                          60
```

Gln Thr Asp Gly Phe Leu Gln Val Gln Tyr Ala Ile Pro Ser Lys Gln

Pro Glu Phe His Ile Glu Ile Leu Ser Ile

<210> 222 <211> 346 <212> PRT <213> Homo sapiens <220> <221> SIGNAL <222> -19..-1

<222> -19..-1 <400> 222 Met Ala Met Ala Gln Lys Leu Ser His Leu Leu Pro Ser Leu Arg Gln -15 -10 Val Ile Gln Glu Pro Gln Leu Ser Leu Gln Pro Glu Pro Val Phe Thr Val Asp Arg Ala Glu Val Pro Pro Leu Phe Trp Lys Pro Tyr Ile Tyr 20 Ala Gly Tyr Arg Pro Leu His Gln Thr Trp Arg Phe Tyr Phe Arg Thr 35 40 Leu Phe Gln Gln His Asn Glu Ala Val Asn Val Trp Thr His Leu Leu 55 Ala Ala Leu Val Leu Leu Arg Leu Ala Leu Phe Val Glu Thr Val Asp Phe Trp Gly Asp Pro His Ala Leu Pro Leu Phe Ile Ile Val Leu Ala Ser Phe Thr Tyr Leu Ser Leu Ser Ala Leu Ala His Leu Leu Gln 100 105 Ala Lys Ser Glu Phe Trp His Tyr Ser Phe Phe Phe Leu Asp Tyr Val 115 120 Gly Val Ala Val Tyr Gln Phe Gly Ser Ala Leu Ala His Phe Tyr Tyr 130 135 Ala Ile Glu Pro Ala Trp His Ala Gln Val Gln Ala Val Phe Leu Pro Met Ala Ala Phe Leu Ala Trp Leu Ser Cys Ile Gly Ser Cys Tyr Asn 165 Lys Tyr Ile Gln Lys Pro Gly Leu Leu Gly Arg Thr Cys Gln Glu Val 180 Pro Ser Val Leu Ala Tyr Ala Leu Asp Ile Ser Pro Val Val His Arg 195 200 Ile Phe Val Ser Ser Asp Pro Thr Thr Asp Asp Pro Ala Leu Leu Tyr 215 210 His Lys Cys Gln Val Val Phe Phe Leu Leu Ala Ala Phe Phe Ser

Thr Phe Met Pro Glu Arg Trp Phe Pro Gly Ser Cys His Val Phe Gly 240 245 250

Gln Gly His Gln Leu Phe His Ile Phe Leu Val Leu Cys Thr Leu Ala

Gln Leu Glu Ala Val Ala Leu Asp Tyr Glu Ala Arg Arg Pro Ile Tyr

<210> 223

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<211> 210
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -20..-1
<400> 223
Met Asp Asn Arg Phe Ala Thr Ala Phe Val Ile Ala Cys Val Leu Ser
                    -15
                                        -10
Leu Ile Ser Thr Ile Tyr Met Ala Ala Ser Ile Gly Thr Asp Phe Trp
                1
Tyr Glu Tyr Arg Ser Pro Val Gln Glu Asn Ser Ser Asp Leu Asn Lys
                           20
        15
Ser Ile Trp Asp Glu Phe Ile Ser Asp Glu Ala Asp Glu Lys Thr Tyr
Asn Asp Ala Leu Phe Arg Tyr Asn Gly Thr Val Gly Leu Trp Arg Arg
                    50
Cys Ile Thr Ile Pro Lys Asn Met His Trp Tyr Ser Pro Pro Glu Arg
                                    70
Thr Glu Ser Phe Asp Val Val Thr Lys Cys Val Ser Phe Thr Leu Thr
                                85
Glu Gln Phe Met Glu Lys Phe Val Asp Pro Gly Asn His Asn Ser Gly
                            100
Ile Asp Leu Leu Arg Thr Tyr Leu Trp Arg Cys Gln Phe Leu Leu Pro
                        115
Phe Val Ser Leu Gly Leu Met Cys Phe Gly Ala Leu Ile Gly Leu Cys
                                        135
                   130
Ala Cys Ile Cys Arg Ser Leu Tyr Pro Thr Ile Ala Thr Gly Ile Leu
                                   150
                145
His Leu Leu Ala Val Thr Lys Glu Ser Met Leu Pro Ala Gly Ala Glu
                                                    170
                                165
Ser Lys His Thr Ala Thr Pro Ala His Ala Cys Val Gln Thr Gly Lys
                            180
Pro Lys
    190
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<210> 224
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<212> PRT
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                                           -30
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                                      -15
Ile Gly Phe Leu Gly Tyr Cys Ser Gly Leu Ile Asp Asn Leu Ile Arg
               -5
Arg Arg Pro Ile Ala Thr Ala Gly Leu His Arg Gln Leu Leu Tyr Ile
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Thr Ala Phe Phe Leu Leu Asp Ile Ile Leu
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Asn Leu Leu Glu Val Cys Lys Lys
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Ile Leu Ala Lys Lys Lys 60

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Arg Gly Glu Met Cys Phe Phe Asp Ser Glu Asp Pro Ala Asn Ser Leu
                      25
Arg Gly Glu Pro Asn Phe Leu Pro Val Thr Glu Glu Ala Asp Ile
                  40
                                      45
Arg Glu Asp Asp Asn Ile Ala Ile Ile Asp Val Pro Val Pro Ser Phe
                                  60
Ser Asp Ser Asp Pro Ala Ala Ile Ile His Asp Phe Glu Lys Gly Met
                              75
Thr Ala Tyr Leu Asp Leu Leu Gly Ile Cys Tyr Leu Met Pro Leu
                   . 90
Asn Thr Ser Ile Val Met Pro Pro Lys Asn Leu Val Glu Leu Phe Gly
                      105
Lys Leu Ala Ser Gly Arg Tyr Leu Pro Gln Thr Tyr Val Val Arg Glu
                  120
                                     125
Asp Leu Val Ala Val Glu Glu Ile Arg Asp Val Ser Asn Leu Gly Ile
               135
                                 140
Phe Ile Tyr Gln Leu Cys Asn Asn Arg Lys Ser Phe Arg Leu Arg Arg
                              155
           150
Arg Asp Leu Leu Gly Phe Asn Lys Arg Ala Ile Asp Lys Cys Trp
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Lys Ile Arg His Phe Pro Asn Glu Phe Ile Val Glu Thr Lys Ile Cys
Gln Glu
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 Gly
 Cys
 Val
 Phe
 Gln
 Ser
 Thr
 Glu
 Asp
 Lys
 Cys
 Ile
 Phe
 Lys
 Ile

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 5
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 7
 10
 10
 15
 15

 Asp
 Trp
 Thr
 Leu
 Ser
 Pro
 Glu
 His
 Ala
 Lys
 Asp
 Glu
 Trp
 Val
 Leu
 Ser
 Val
 Pro
 Ile
 Gly
 Arg
 Phe
 Gln
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 Leu<

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 Phe Glu Val Asn
 Pro Trp Lys Arg Glu Val Lys Leu Leu Leu Ser Ser

 120
 125
 130

 Glu Thr Pro Ile Glu Gly Lys Asn Met Ser Phe Val Asn Asp Leu Thr
 135
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 145

Lys Thr Arg Asp Asp Glu Pro Val Cys Gly Arg Pro Leu Gly Ile Arg

Ala Gly Pro Asn Gly Thr Leu Phe Val Ala Asp Ala Cys Lys Gly Leu

110

Val Ser Gln Asp Gly Arg Lys Ile Tyr Phe Thr Asp Ser Ser Lys 155 Trp Gln Arg Arg Asp Tyr Leu Leu Leu Val Met Glu Gly Thr Asp Asp 170 175 Gly Arg Leu Leu Glu Tyr Asp Thr Val Thr Arg Glu Val Lys Val Leu 185 190 Leu Asp Gln Leu Arg Phe Pro Asn Gly Val Gln Leu Ser Pro Ala Glu 200 205 Asp Phe Val Leu Val Ala Glu Thr Thr Met Ala Arg Ile Arg Arg Val 220 Tyr Val Ser Gly Leu Met Lys Gly Gly Ala Asp Leu Phe Val Glu Asn 235 Met Pro Gly Phe Pro Asp Asn Ile Arg Pro Ser Ser Ser Gly Gly Tyr 250 255 Trp Val Gly Met Ser Thr Ile Arg Pro Asn Pro Gly Phe Ser Met Leu 265 270 Asp Phe Leu Ser Glu Arg Pro Trp Ile Lys Arg Met Ile Phe Lys Ala Lys Lys Lys

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-15 -10 -5

Leu Arg Glu Phe Ser Gln Ile Arg Tyr Asp Ala Val Lys Ser Lys Met
1 5 10 15

Asp Pro Glu Leu Glu Lys Lys Leu Lys Glu Asn Lys Ile Ser Leu Glu 20 25 30

Ser Glu Tyr Glu Lys Ile Lys Asp Ser Lys Phe Asp Asp Trp Lys Asn 35 40 45

Ile Arg Gly Pro Arg Pro Trp Glu Asp Pro Asp Leu Leu Gln Gly Arg 50 55 60

Asn Pro Glu Ser Leu Lys Thr Lys Thr Thr

<210> 237

<211> 42

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -19..-1

<400> 237

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Gln Leu Ser Asp Lys Val His Asn Asp Ile
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                                        -10
Leu Ile Ser Thr Ile Tyr Met Ala Ala Ser Ile Gly Thr Asp Phe Trp
                                                    10
               ٦
Tyr Glu Tyr Arg Ser Pro Val Gln Glu Asn Ser Ser Asp Leu Asn Lys
                            20
Ser Ile Trp Asp Glu Phe Ile Ser Asp Glu Ala Asp Glu Lys Thr Tyr
                       35
                                            40
Asn Asp Ala Leu Phe Arg Tyr Asn Gly Thr Val Gly Leu Trp Gly Arg
                    50
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Cys Ile Thr Ile Pro Lys Asn Met His Trp Tyr Ser Pro Pro Glu Arg
                                    70
Thr Gly Ile Ser Leu Ile Leu Thr Ser Val Phe Phe Thr Trp Leu Ile
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Ile Asp Lys Thr Thr
        95
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Gln His Xaa Xaa Ala Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile
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Leu Thr Ile Leu Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe
Leu His Glu Thr Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu
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Ile Ser Arg Tyr Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val
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Cys Asp Cys Val Lys Leu Thr Phe Ser Pro Pro Thr Leu Leu Val Asn
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Val Thr Asp Gln Val Tyr Glu Tyr Lys Tyr Lys Arg Glu Ile Ser Gln

His Asn Ile Asn Pro His Gln Gly Asn Ala Ile Leu Glu Lys Met Thr

Phe Asp Pro Glu Ile Phe Phe Asn Val Leu Pro Pro Ile Ile Phe

65

80

70

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His Ala Gly Tyr Ser Leu Lys Lys Arg His Phe Phe Gln Asn Leu Gly
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Ile Gly
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 Met Cys Ala Leu Val Thr Phe Phe Phe Ile Leu Leu Leu Leu Ile Phe Ile 10

 Ala Glu Val Ala Ala Ala Val Val Ala Leu Val Tyr Thr Thr Met Ala 25

 Glu His Phe Leu Thr Leu Leu Val Val Val Pro Ala Ile Lys Lys Asp Tyr 40

 Gly Ser Gln Glu Asp Phe Thr Gln Val Trp Asn Thr Thr Met Lys Gly 55

 Leu Lys Cys Arg Gly Phe Thr Asn Tyr Thr Asp Phe Glu Asp Ser Pro 70

 Tyr Phe Lys Met His Lys Pro Val Thr Met Lys Lys Lys Lys Lys

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-15 Ile Glu Gly Gly Val Val Phe Tyr Gln Leu Tyr Ser Leu Leu Arg Ser-Glu Lys Trp Asn His Thr Leu Ser Met Ala Leu Ile Leu Phe Cys Asn 20 Tyr Tyr Val Leu Phe Lys Leu Leu Arg Asp Arg Ile Val Leu Gly Arg 40 Ala Tyr Ser Tyr Pro Leu Asn Ser Tyr Glu Leu Lys Ala Asn 50

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896

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	8 0
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	04
att gca att gtg tta ctt tta ttt gga tgaatwtcat tggagaaaat 35 Ile Ala Ile Val Leu Leu Leu Phe Gly 60 65	54
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884

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897

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                                                                  109
              Met Lys Ala Leu Cys Leu Leu Leu Pro Val Leu
                          -15
                                             -10
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Gly Leu Leu Val Ser Ser Lys Thr Leu Cys Ser Met Glu Glu Ala Ile
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aat gag agg atc cag gag gtc gcc ggc tcc cta ata ttt agg gca ata
Asn Glu Arg Ile Gln Glu Val Ala Gly Ser Leu Ile Phe Arg Ala Ile
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age age att gge ega ggg age gag age gte ace tee agg ggg gae etg
Ser Ser Ile Gly Arg Gly Ser Glu Ser Val Thr Ser Arg Gly Asp Leu
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                                                                  301
get act tgc ccc cga ggc ttc gcc gtc acc ggc tgc act tgt ggc tcc
Ala Thr Cys Pro Arg Gly Phe Ala Val Thr Gly Cys Thr Cys Gly Ser
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gcc tgt ggc tcg tgg gat gtg cgc gcc gag acc aca tgt cac tgc cag
                                                                   349
Ala Cys Gly Ser Trp Asp Val Arg Ala Glu Thr Thr Cys His Cys Gln
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Cys Ala Gly Met Asp Trp Thr Gly Ala Arg Cys Cys Arg Val Gln Pro
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271

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ttc ctg wgt cta atg acc ctg aca acc cat gtt cac tca agt gcc aag 164

Phe Leu Xaa Leu Met Thr Leu Thr Thr His Val His Ser Ser Ala Lys -10 -5 1

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Pro Asn Glu Gln Pro Trp Leu Leu Asn 5 10

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> 331 391

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gga cag ac Gly Gln Th			sp Thr										152
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cta gtg ac Leu Val Th													344
act gat ga Thr Asp As 80		Thr T											392

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atg acc cct tct tct atg atg aac aca ccc tcc gga aac sgg ggc tgt Met Thr Pro Ser Ser Met Met Asn Thr Pro Ser Gly Asn Xaa Gly Cys 115 120 125	488											
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Arg Pro Val Leu Gln Asn Leu Leu Gln Ser Pro Gly Leu Thr Trp Ser  1 5 10 15												
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CCa ggg cta akt ggg gcc act cca gcc agg agc cct cag ggc aag gag Pro Gly Leu Xaa Gly Ala Thr Pro Ala Arg Ser Pro Gln Gly Lys Glu 35 40 45	245											
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acg ggc ctc cat tcc tgc ara gat ggt atg gct tct ctt gaa ggg acg	437										
Thr Gly Leu His Ser Cys Xaa Asp Gly Met Ala Ser Leu Glu Gly Thr 100 105 110	•••										
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Pro Ala Ser Val Leu Ala Asp Ala Cys Pro Gly Phe His Asp Val Xaa											
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80 85 90												
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cgt Arg	agc Ser	Ile	999 Gly ,115	gga	wca Xaa	ccg Pro	ccc Pro	amc Xaa 120	ctg	ggt Gly	gcc Ala	ttg Leu	gac Asp 125	ctg Leu	ctg Leu	601
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Leu Lys	Leu	Lys	Lys	Pro	Pro	Trp	Leu	His	Met	Pro	Ser	Ala	Met	Thr	
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Val Tyr	Ala	Leu	Val	Val	Val	Ser	Tyr	Phe	Leu	Ile	Thr	Gly	Gly	Ile	
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Ile Tyr	Asp	Val	Ile	Val	Glu	Pro	Pro	Ser	Val	Gly	Ser	Met	Thr	Asp	
	-50					-45					-40				
gaa cat	999	cat	cag	agg	cca	gta	gct	ttc	ttg	gcc	tac	aga	gta	aat	298
Glu His	Gly	His	Gln	Arg	Pro	Val	Ala	Phe	Leu	Ala	Tyr	Arg	Val	Asn	
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Gly Gln	Tyr	Ile	Met	Glu	Gly	Leu	Ala	Ser	Ser	Phe	Leu	Phe	Thr	Met	
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Gly Gly	Leu	Gly	Phe	Ile	Ile	Leu	Asp	Gly	Ser	Asn	Ala	Pro	Asn	Ile	
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Pro Lys															
	15		_			20				-	25		-		
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Leu Xaa															
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Gly Tyr								_		_				_	
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Leu Ser Wal Thr Gln Pro Trp Tyr Leu Glu Val Asp Tyr Thr His Glu

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															gag	194
Ala	Val	Thr	Ile	Lys 20	Cys	Thr	Phe	Ser	Ala 25	Thr	Gly	Cys	Pro	Ser 30	Glu	
		aca														242
Gln	Pro	Thr	Cys 35	Leu	Trp	Phe	Arg	Tyr 40	Gly	Ala	His	Gln	Pro 45	Glu	Asn	
_	_	ttg	_		_		_		_	_	_					290
	-	Leu 50	_	-	-	-	55				-	60			_	
		ctc														338
Glu	Ala 65	Leu	Lys	Glu	Asn	Gln 70	Val	Ser	Leu	Thr	Val 75	Asn	Arg	Val	Thr	
		gac	_	_				_			•			_		386
Ser 80	Asn	Asp	Ser	Ala	Ile 85	Tyr	Ile	Сув	Gly	Ile 90	Ala	Phe	Pro	Ser	Val 95	
	~~~	gcg	202	act		cad	202	aas	aaa		200	202	cta	ata		434
		Ala														323
			5	100	-,-			1	105	,				110		
aga	gaa	att	aag	ctg	ctc	agc	aag	gaa	ctg	cgg	agc	ttc	ctg	aca	gct	482
Arg	Glu	Ile	Lys 115	Leu	Leu	Ser	Lys	Glu 120	Leu	Arg	Ser	Phe	Leu 125	Thr	Ala	
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Leu	Val	Ser 130	Leu	Leu	Ser	Val	Tyr 135	Val	Thr	Gly	Val	Cys 140	Val	Ala	Phe	
ata	ctc	ctc	tcc	aaa	tca	aaa	tcc	aac	cct	cta	aga	aac	aaa	gaa	ata	578
Ile	Leu 145	Leu	Ser	Lys	Ser	Lys 150	Ser	Asn	Pro	Leu	Arg 155	Asn	Lys	Glu	Ile	
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_	Glu	Asp	Ser	Gln	_	Lys	Lys	Ser	Ala	_	Arg	Ile	Phe	Gln		
160					165					170					175	674
	_	caa Gln	_				_	_			_			_		674
116	MIG	GIII	Giu	180	ıyı	итэ	пуъ	Arg	185	Val	GIU	TIII	ASII	190	GIII	
tct	gag	aaa	gat		aac	act	tat	gaa		aga	aga	gta	ctt		aac	722
Ser	Glu	Lys	Asp 195	Asn	Asn	Thr	Tyr	Glu 200	Asn	Arg	Arg	Val	Leu 205	Ser	Asn	
tat	gaa	agg	cca	taga	aaac	gtt 1	ttaat	ttt	ca at	tgaag	gtcad	tga	aaaat	tcca		774
Tyr	Glu	Arg 210	Pro													
act	ccag	gag	ctate	ggca	gt gi	taai	tgaad	ata	atato	catc	aggt	ctta	aaa a	aaaa	ataaa	834
					_		_								tataac	894
taa	takt	cat 1	tacc	aaaa	ta c	caaa	accc	a aca	aaaa	tgca	act	gaaaa	aat a	accti	tccaaa	954
ttt	gcca	aaa a	aaaa	aaw												971

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-30 -25 -20 tgg ttg gaa gta gaa tgt tcg ctg atg acc tgc aca acc tta ata aac Trp Leu Glu Val Glu Cys Ser Leu Met Thr Cys Thr Thr Leu Ile Asn	458
gca tcc gct atc tct aca aac act tta acc gac atg gga agt ttc gat Ala Ser Ala Ile Ser Thr Asn Thr Leu Thr Asp Met Gly Ser Phe Asp  1 5 10	506
aga aga gaa agc tgagaacttc ggaaaaggct catctgtcac cctggaraag Arg Arg Glu Ser	558
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ggc ttg cag atg ttc att cag agg aag ttt cca tac cct ttg cag tgg Gly Leu Gln Met Phe Ile Gln Arg Lys Phe Pro Tyr Pro Leu Gln Trp -25 -20 -15 -10	166
agc ctc cta gtg gcc gtg gtt gca ggc tct gtg gtc agc tac ggg gtg Ser Leu Leu Val Ala Val Val Ala Gly Ser Val Val Ser Tyr Gly Val	214
acg aga gtg gag tcg gag aaa tgc aac aac ctc tgg ctc ttc ctg gag	262

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Thr Arg Val Glu Ser Glu Lys Cys Asn Asn Leu Trp Leu Phe Leu Glu 10 15 20	
acc gga cag ctc ccc aaa gac agg agc aca gat cag ara agc Thr Gly Gln Leu Pro Lys Asp Arg Ser Thr Asp Gln Xaa Ser 25 30 35	304
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ctgtccgcat gtgtggccag gcctgacaaa cmcctgcaga tggctgctgc cccaacctgg	484
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Met Thr Cys Trp Met Leu Pro Pro Ile Ser Phe	233
-20 -15	
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Leu Ser Tyr Leu Pro Leu Trp Leu Gly Pro Ile Trp Pro Cys Ser Gly -10 -5 1	
-10 -5 1 tet acc ett ggg aag eet gat eee ggt gtg tgg eee age ttg tte agg	329
Ser Thr Leu Gly Lys Pro Asp Pro Gly Val Trp Pro Ser Leu Phe Arg	323
5 10 15	
ccc tgg gat gct gca tct cca ggc aac tat gca ctt tcc cgg gga rar	377
Pro Trp Asp Ala Ala Ser Pro Gly Asn Tyr Ala Leu Ser Arg Gly Xaa 20 35	
20 25 30 35 aac cak tat gav aak tgg ggg cag ggc aca cat tca tct ttg	419
Asn Xaa Tyr Xaa Xaa Trp Gly Gln Gly Thr His Ser Ser Leu	140
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gacctgottt gocattotoc tggtgccgct gctgctccct gtttctggag ctggatgttc	539
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                                  -10
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Thr Ser Ser His Ala Ser Ser Leu His Leu Pro Pro Ser Cys Thr Arg
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Leu Thr Leu Thr Gln Thr Leu Arg Thr Gly Met His Leu Ser Arg Ala
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Leu Gln Gly Thr Leu Thr Arg Leu Gln Ser Thr Pro Ala
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Ile Ser Ile Phe Pro Thr Met Met Val Cys Met Met Ala Trp Arg Pro
                                         -50
                    -55
att cag gca ctt atg gcc att tca gcc act ttc aag atg tta gaa agt
                                                                      148
Ile Gln Ala Leu Met Ala Ile Ser Ala Thr Phe Lys Met Leu Glu Ser
                                     -35
                -40
tca agc cag aag ttt ctt cag ggt ttg gtc tat ctc att ggg aac ctg
                                                                      196
Ser Ser Gln Lys Phe Leu Gln Gly Leu Val Tyr Leu Ile Gly Asn Leu
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            -25
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Met Gly Leu Ala Leu Ala Val Tyr Lys Cys Gln Ser Met Gly Leu Leu
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Pro Thr His Ala Ser Asp Trp Leu Ala Phe Ile Glu Pro Pro Glu Arg
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Met Glu Ser Val Val Glu Asp Cys Phe Cys Glu His Glu Lys Ala Ala
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                                                                       388
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Pro Gly Pro Tyr Val Phe Gly Ser Tyr Leu His Pro Ser Leu Ser Pro
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gtg gct cct cag cat act ctt aaa cta act tat gtt aaa aaa aac
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Val Ala Pro Gln His Thr Leu Lys Leu Ile Thr Tyr Val Lys Lys Asn
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Gln Lys Thr Leu Phe Ser Met Val Gly
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                                                                       663
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caggagetee gggaggeagg geeggeeeca egteetetge geaceaceet gagttggate
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ctctgtgcgc cacccctgag ttggatccag ggctagctgc tgttgacctc cccactccca
                                                                      240
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Val Leu Met Gly Leu Pro Leu Ala Gln Ala Leu Asp Cys His Val Cys
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Ala Tyr Asn Gly Asp Asn Cys Phe Asn Pro Met Arg Cys Pro Ala Met
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gtt gcc tac tgc atg acc acg cgc acc tac tac acc ccc acc agg atg
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Val Ala Tyr Cys Met Thr Thr Arg Thr Tyr Tyr Thr Pro Thr Arg Met
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Lys Val Ser Lys Ser Cys Val Pro Arg Cys Phe Glu Xaa Cys Val
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getayaccet ckcacccake teaccetgee teacceteea cactecetge gaccteetea
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WO 99/31236 -186- PCT/IB98/02122

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tgt ggg tct ctc cta cct ggg tta tgg cag cac ctc aca gcc aat cac Cys Gly Ser Leu Pro Gly Leu Trp Gln His Leu Thr Ala Asn His

1

-5

-10

326

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tgg cct cca ttc tcc sct ttc ctc tgt aca gtt tgc tct ggt tcc tca  Trp Pro Pro Phe Ser Xaa Phe Leu Cys Thr Val Cys Ser Gly Ser Ser  10 15 20	326
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tetgetecce ttggtttect cetgtraart aaateteact gaccettgat geasetecaa	531
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gcaaagccaa cectcacege tggteggtgg gccatace atg gga aag gga cat cag  Met Gly Lys Gly His Gln  -25	296
cgg ccc tgg tgg aag gtg ctg ccc ctc agc tgc ttc ctc gtg gcg ctg Arg Pro Trp Trp Lys Val Leu Pro Leu Ser Cys Phe Leu Val Ala Leu -20 -15 -10	344
Arg Pro Trp Trp Lys Val Leu Pro Leu Ser Cys Phe Leu Val Ala Leu -20 -15 -10 atc atc tgg tgc tac ctg agg gag gag agc gag gcg gac cag tgg ttg	344
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Arg Gln Val Trp Gly Glu Val Pro Glu Pro Ser Asp Arg Ser Glu Glu	
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ctg gtg cac ttc ata gct ttt tat cta ata ttt ctc aac atg cct gga Leu Val His Phe Ile Ala Phe Tyr Leu Ile Phe Leu Asn Met Pro Gly 55 60 65	290
gat gcc ccg att gct cct gtt aaa gga act gac agc agt gct tac atc Asp Ala Pro Ile Ala Pro Val Lys Gly Thr Asp Ser Ser Ala Tyr Ile 70 75 80	338
aaa too ago aaa raa ttt goo att oto tgo akt ttt otg tkg ggo ott Lys Ser Ser Lys Xaa Phe Ala Ile Leu Cys Xaa Phe Leu Xaa Gly Leu	386

90 95	
85 90 95  qqa aac agc tgc ttt aat acc cas ctg ctt akt atc tkg ggc ttt ctg	434
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100 105 110	
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Tyr Ser Glu Xaa Ser Ala Pro Xaa Phe Ala Ile Phe Asn Phe Val Gln	
115 120 125 130 tet att tge gea gee gtg gea ttt tte tae age aac tae ett ete ett	530
Ser Ile Cys Ala Ala Val Ala Phe Phe Tyr Ser Asn Tyr Leu Leu	
135 140 145	
cac tgg caa ctc ctg gtc atg gtk atw ttt ggg ttt ttk gga aca att	578
His Trp Gln Leu Leu Val Met Val Ile Phe Gly Phe Xaa Gly Thr Ile	
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Ser Asp Tyr Arg Ser Ile 180	
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Asn Lys Ser Tyr Lys Asn Lys Asp Ser Val Arg Ile Tyr Leu Ser Leu	
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Cys Thr Val Ser Ile Lys Phe Thr Tyr Phe His Asp Ile Gln Thr Asn	
-55 -50 -45	258
tgt ctt aca aca tgg aaa cat tcg aga tgc aga ttt tat tgg gca ttt Cys Leu Thr Thr Trp Lys His Ser Arg Cys Arg Phe Tyr Trp Ala Phe	
-40 -35 -30 -25	
ggt ggt tee att tta cag cae tea gtg gat eee ett gtt ttg tte eta	306
Gly Gly Ser Ile Leu Gln His Ser Val Asp Pro Leu Val Leu Phe Leu	
GIV GIV SET THE LEG GIT HIS SET VAL ASP FLO DEG VAL DEC THE DEC	

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Ser Leu Ala Leu Leu Val Thr Pro Thr Ser Thr Pro Ser Ala Lys Ile	
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car ago ott caa att gac otc oot gga ggo tgg agg otg goo act gac	402
Gln Ser Leu Gln Ile Asp Leu Pro Gly Gly Trp Arg Leu Ala Thr Asp	
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His Gln Leu	559
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	Ala Lei													
	-10				-5					1				
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98

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Lys	Ile	Asn	Asp		Thr	Gin	GIU	PTO	Val 40	Asn	Cys	Thr	ASII	Tyr 45	TIII		
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gct	cat	gtt	Com	cgt.	Dho	Dro	yca Nla	Dro	Acn	Tla	Thr	Cyc	Lve	gat Asp	Ser	250	
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ccc	ata		tac	cqa	aat	qta	aat	ggc	tat	tcc	tac	aat	gag	cag	tcg	386	,
Pro	Ile	Ser	Cys	Arg'	Asn	Val	Asn	Gly	Tyr	Ser	Tyr	Asn	Glu	Gln	Ser		
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cat	gtc	tct	ttt	tct	tgg	atg	gtt	<b>9</b> 99	agc	aga	tcg	att	tta	cct	tgg	434	:
His	Val	Ser	Phe	Ser	Trp	Met	Val	Gly	Ser	Arg	Ser	Ile	Leu	Pro			
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Ile	Pro	Cys	Phe		Phe	Val	Lys	Xaa		His	Суѕ	Arg	Val	Xaa	Trp		
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					tga	tttc	ату о	ccca	CCCC	aa t	gcag	actg	c cg	gacci	LLCa	537	
Asn	Trp	Glu	Pro	Asn													
+	~~~~	<b></b>	130	a=++:	a t- a	astt:	actai	t aa	2200	agac	tta	caag	act (	gagta	attact	: 597	,
aatt	yyaa	yea :	great ttam	2222	מנ מי	raati	tatai	t cc	ataa	atat	ttt	ttaa	aag	aaac	agattt		
gagg	ctc	ott i	gatti	ttaai	ta or	agaa	cttc	t ag	tata	taga	ttt	aaaq	att	tctc	ttttt	: 717	,
atto	cata	tac	catti	ttate	ga g	ttct	gtata	a at	tttt	tata	qtt	tttg	ttt	tgtt	gagtta	a 777	,
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aaaa	ggca	aqt	tgcc	actt	gt c	attt	ttgti	t tc	tgaa	aaat	aaa	agta	taa	ctta	ttcaca	a 957	t
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	-			-25					-20					-15		
atg Met	ttg Leu	ata Ile	Met	ctc Leu	gga Gly	ata Ile	ttt Phe	Phe	aat Asn	gtc Val	cat His	tcc Ser	Ala	gtg Val	ttg Leu	209
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att Ile	gag Glu 5	gac Asp	gtt Val	Pro	Phe	acg Thr 10	gag Glu	Lys	Asp	Phe	gag Glu 15	Asn	Gly	ccc Pro	Gln	231
226		tac	220	ctt	tac		caa	ktc	age	tac		tat	ttc	atc	act	305
A a c	Tle	Tyr	Acn	Len	Tvr	Xaa	Gln	Xaa	Ser	Tvr	Asn	Cvs	Phe	Ile	Āla	
20	116	TYT	You	шец	25		0111			30		-1-			35	
	~~~	a++	+ > 0	ctc		ctc	gga	ggc	ttc	_	ttc	tac	caa	ktt	caa	353
gca	234	Len	Tur	Leu	T.eu	Len	Glv	Glv	Phe	Ser	Phe	Cvs	Gln	Xaa	Arq	
Ald	GIY	Den	TAT	40	шец	шси	ULY	ردن	45			<b>-</b> 1		50	5	
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Leu	Asn	Lys	Arg 55	Lys	Glu	Tyr	Met	Val 60	Arg	cag	9900		3030	5000		
cca	ėt cc	agc		cata	ta t	ttaa	arac		ctac	accq	tkt	cacc	cag	gtcg	cgtccc	463
acc	ctta	cca .	acac	cata	ta t	ggga	ctaa	a tt	tece	aaac	rar	arac	tga	atco	cttctc	523
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COL	cctt	cac	tata	toco	at a	tata	aata	a ag	agaa	tcta	ctc	tctt	caa	aaaa	aaaaaa	643
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		-														
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45 50 55 tgaaaagaaa gtgttttatt ttcaaatctg gtccatattt acattctagt tcagagccaa gccttaaact gtacagaatt tccactgtaa ttaaaactat ttagtgttag ttataaatag	601 661
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ttt aat tgg ttc gac tgc ctt ctc cac aat ttg ggc gag aat ttc ctt Phe Asn Trp Phe Asp Cys Leu Leu His Asn Leu Gly Glu Asn Phe Leu	150
-25 -20 -15 -10  ago oft oft ago aaa agt tot tot gog gad oog tot gog toa act too  Ser Leu Leu Ser Lys Ser Cys Ser Ala Asp Pro Ser Gly Ser Thr Phe	198
-5 1 5 atg agg gac att gag aca aac aaa tgaaatatgg gttaaagtac tctgagcagc	252
Met Arg Asp Ile Glu Thr Asn Lys 10 15	
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ctaactacat gagtgagacc agttgacaac acatggagca racatgagct gttctcagtg artcctacac aaattcctga ctcacaacac tgtgagcaat aaaatggttg ttattttaag	492 552
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                                                                      117
ccc gag gct gtg gaa caa tca gcc cat ctc ttt gtg acc tgg agc agt
                                                                      165
Pro Glu Ala Val Glu Gln Ser Ala His Leu Phe Val Thr Trp Ser Ser
                         -15
                                             -10
    -20
cag agg gcc ctc agt cac ccc gcc cca ttc ctc acc ara raa aar aat
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Gln Arg Ala Leu Ser His Pro Ala Pro Phe Leu Thr Xaa Xaa Lys Asn
                                                                       261
cca ttt cta tgg aag ctc tgacgtaact tcagtgtttt ctacaatact
Pro Phe Leu Trp Lys Leu
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agocttagtt toccatggod otgaaacaca cacatttood cottoottto ccagaagoda	180
ctggccccc atagcaccca gtgcatcctt tttacaagtg gaagaactag g atg gct Met Ala	237
tto caa agt ott ota gaa atg aag tto ttt oto tgt goa got tto coo	285
Phe Gln Ser Leu Leu Glu Met Lys Phe Phe Leu Cys Ala Ala Phe Pro -20 -15 -10	
ctt gga gca gga gtg aag atg ttt cat tat ctt ggg cct ggg aaa cca	333
Leu Gly Ala Gly Val Lys Met Phe His Tyr Leu Gly Pro Gly Lys Pro	
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Leu Xaa Gln Ala Ser Pro Ser Pro His Pro His Arg Xaa Arg Ile Trp	
15 20 25	
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Pro	
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ctcatttttt tottotggto agttttttta agggggggtg ttgtggtttt ttgtttttgt	554
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agagcaagtg gaatctctaa ga atg gct tcc agc cac tgg aat gaa acc act	172
Met Ala Ser Ser His Trp Asn Glu Thr Thr	
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Phe His Asp Asn Trp Asn Thr Ala Cys Phe Val Ile Leu Leu Phe	
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ata ttt aca gtg gta tct tta gtg gtg ctg gct ttc ctt tat gaa gtg	316
Ile Phe Thr Val Val Ser Leu Val Val Leu Ala Phe Leu Tyr Glu Val	
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Leu Xaa Xaa Cys Cys Cys Val Lys Asn Lys Thr Val Lys Asp Leu Lys	

15 20 25	
agt gaa ccc aac cct ctt ara akt atg atg gac aac atc aga aaa cgt	412
Ser Glu Pro Asn Pro Leu Xaa Xaa Met Met Asp Asn Ile Arg Lys Arg	
30 35 40	
gaa act gaa gtg gtc taacactcta taraaaatga acaaaatctc tgaaagcagc	467
Glu Thr Glu Val	
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tcaacctctt ctgaraaaaa aaatatattc tgaggccaac tgttgctaca aaacaaattc	527
tgactgaatg gttaaaacat ttctagtara aggggaaaaa aaakttaaac atgcactgtt	587
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cactteetga gtgageteae ttacetteee tgaatggtga gge atg gat gaa tat	295
Met Asp Glu Tyr	
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tee tgg tgg tge cae gtg tta gag gtg gta aag ggt caa atg ttt act	343
Ser Trp Trp Cys His Val Leu Glu Val Val Lys Gly Gln Met Phe Thr	
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Phe Ile Asn Ile Thr Leu Trp Leu Gly Ser Leu Cys Gln Arg Phe Phe	
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Tyr Ala Ser Gly Thr Tyr Phe Leu Ile Tyr Ile Ser Thr Val Thr Pro	
5 10 15 20	
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Ser Trp Arg Leu Cys Leu Val Ser	
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                                                                      107
   Met Tyr Val Trp Pro Cys Ala Val Val Leu Ala Gln Tyr Leu Trp
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                                                                      155
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Phe His Arg Arg Ser Leu Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala
        -65
                            -60
gga gtg agc ctt cca gga att ttg gct gcc aaa tgt ggt gca gaa gta
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Gly Val Ser Leu Pro Gly Ile Leu Ala Ala Lys Cys Gly Ala Glu Val
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Ile Leu Ser Asp Ser Ser Glu Leu Pro His Cys Leu Glu Val Cys Arg
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                    -30
                                         -25
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                                     -10
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Thr Trp Gly His Ile Ser Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp
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Ile Ile Leu Ala Ser Asp Val Phe Phe Glu Pro Glu Xaa Phe Glu Asp
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att ttg gct aca ata tat ttt ttg atg cac aar aat ccc aag gtc caa
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Ile Leu Ala Thr Ile Tyr Phe Leu Met His Lys Asn Pro Lys Val Gln
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                                                                      539
Leu Leu Tyr Lys Trp Asp Met Lys Cys Val His Ile Pro Leu Glu Ser
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Phe Asp Ala Asp Lys Glu Xaa Ile Ala Glu Ser Thr Leu Pro Gly Arg
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His Thr Val Glu Met Leu Val Ile Ser Phe Ala Lys Asp Ser Leu
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812

840 .

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too coc cag goo ctg gag gac tog ggo cog gtg aat ato toa gto toa
                                                                      101
Ser Pro Gln Ala Leu Glu Asp Ser Gly Pro Val Asn Ile Ser Val Ser
        -95
                             -90
ate ace eta ace etg gae eca etg aaa ece tte gga ggg tat tee ege
                                                                      149
Ile Thr Leu Thr Leu Asp Pro Leu Lys Pro Phe Gly Gly Tyr Ser Arg
                        -75
                                             -70
aac gtc acc cat ctg tac tca acc atc tta ggg cat cag att gga ctt
                                                                      197
Asn Val Thr His Leu Tyr Ser Thr Ile Leu Gly His Gln Ile Gly Leu
                    -60
                                         -55
tca ggc agg gaa gcc cac gag gag ata aac atc acc ttc acc ctg cct
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Ser Gly Arg Glu Ala His Glu Glu Ile Asn Ile Thr Phe Thr Leu Pro
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aca gcg tgg agc tca gat gac tgc gcc ctc cac ggt cac tgt gag cag
                                                                      293
Thr Ala Trp Ser Ser Asp Asp Cys Ala Leu His Gly His Cys Glu Gln
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                                 -25
                                                     -20
gtg gta ttc aca gcc tgc atg acc ctc acg gcc agc cct ggg gtg ttc
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Val Val Phe Thr Ala Cys Met Thr Leu Thr Ala Ser Pro Gly Val Phe
                             -10
ecg tea etg tac age cac ege act gtg tte etg aca egt aca gea acg
                                                                      389
Pro Ser Leu Tyr Ser His Arg Thr Val Phe Leu Thr Arg Thr Ala Thr
cca cgc tct ggt aca aga tct tca caa ctg cca gag atg cca aca caa
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Pro Arg Ser Gly Thr Arg Ser Ser Gln Leu Pro Glu Met Pro Thr Gln
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aat acg ccc aaa att aca atc ctt tct ggt gtt ata agg ggg cca ttg
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Asn Thr Pro Lys Ile Thr Ile Leu Ser Gly Val Ile Arg Gly Pro Leu
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gaa aag tot atc atg ott taaatoocaa gottacagtg attgttocag
Glu Lys Ser Ile Met Leu
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cacgtgcttt gccctgggta cagccagagc ccttcaaccc caccttggac ttgaggacct
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1013

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gatgttttca ccaaggtcac aggagcattg cgtcgctgat ggggttgaag tttggtttgg	1073 1133 1193 1253 1313 1344
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caa agg cca gag tca cag gaa gga ctt ctt cca ggg aga tta gtg gtg Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro Gly Arg Leu Val Val -5 1 5 10	96
atg gag agg aga gtt aaa aat gac ctc atg tcc ttc ttg tcc acg gtt Met Glu Arg Arg Val Lys Asn Asp Leu Met Ser Phe Leu Ser Thr Val 15 20 25	144
ttg ttg agt ttt cac tct tct aat gca agg gtc tca cac tgt gaa cca Leu Leu Ser Phe His Ser Ser Asn Ala Arg Val Ser His Cys Glu Pro	192
30 35 40 Ctt agg atg tgatcacttt caggtggcca ggaatgttga atgtctttgg Leu Arg Met 45	241
ctcagttcat ttaaaaaaga tatctatttg aaagttctca rarttgtaca tatgtttcac	301
agtacaggat ctgtacataa aagtttcttt cctaaaccat tcaccaagag ccaatatcta	361
ggcattttct tggtagcaca aattttctta ttgcttaraa aattgtcctc cttgttattt ctgtttgtaa racttaagtg agttaggtct ttaaggaaag caacgctcct ctgaaatgct	421 481
tgtctttttt ctgttgccga aatarctggt cctttttcgg gagttaratg tatarartgt	541
ttgtatgtaa acatttcttg taggcatcac catgaacaaa gatatattt ctatttattt	601
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WO 99/31236

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185 190 195	
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Ser Leu Met Pro Val Leu Gly Xaa Asp Pro Pro Gln Leu Tyr Leu Thr	
200 205 210	
cag ctc arg gag gcc ttt ggg gat ctg gcc ctt ttc ttc tat gac cag	824
Gln Leu Xaa Glu Ala Phe Gly Asp Leu Ala Leu Phe Phe Tyr Asp Gln 215 220 225	
225 230	
cat ggt gga gag gtg att ggt gtc ctc tgg aag ccc acc agc ttc cag	872
His Gly Gly Glu Val Ile Gly Val Leu Trp Lys Pro Thr Ser Phe Gln	
240	
ccg cag ccc ttc aag gcc tcc agc aca aag ggg cgc atg gtg atg tct Pro Gln Pro Phe Lys Ala Ser Ser Thr Lys Gly Arg Met Val Met Ser	920
250 255 260	
cga ggt ggg gag cta gta atg gtg ccc aat gtt gaa gca atc ctg gag	0.50
Arg Gly Glu Leu Val Met Val Pro Asn Val Glu Ala Ile Leu Glu	968
265 270 275	
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Asp Phe Ala Val Leu Gly Glu Gly Leu Val Gln Thr Val Glu Ala Arg	1010
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Ser Glu Arg Trp Thr Val	1001
295 300	
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-5 1 5 10	
aga cac cac ata ctg cag cag ttc cta gtg aga aaa tct gtg cca cta	206 .
Arg His His Ile Leu Gln Gln Phe Leu Val Arg Lys Ser Val Pro Leu	
15 20 25	
gaa aat gct tca ctt cca ttt cct cac ctg ggc agt tct ctg ttt aaa	254
Glu Asn Ala Ser Leu Pro Phe Pro His Leu Gly Ser Ser Leu Phe Lys	
30 35 40	
att gtg ggc tgatttggtc ttcctctcct cctcccactg ttactgccct	2.02
Ile Val Gly	303
45	
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acacaccett cegeceaaat acetetgace ecaaggetgg aatggggetg gtaggarata	423
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Met	11/
-30	
	1.00
CCa act ggc aag cag cta gct gac att ggc tat aag acc ttc tct acc	165
Pro Thr Gly Lys Gln Leu Ala Asp Ile Gly Tyr Lys Thr Phe Ser Thr	
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-10 -5 1	
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Val Tyr His Tyr Phe Gln Trp Arg Arg Ala Glr 5 10	n Arg Gln Ala Ala Glu 15
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Met Lys Lys	Val Leu Leu Ile
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Thr Ala Ile Leu Ala Val Ala Val Gly Phe Pro	Val Ser Gln Asp Gln
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Glu Arg Glu Lys Arg Ser Ile Ser Asp Ser Asp	Glu Leu Ala Ser Gly
10 15	20
wit tit gtg ttc cct tac cca tat cca tit cgc	cca ctt cca cca att 257
Xaa Phe Val Phe Pro Tyr Pro Tyr Pro Phe Arg	Pro Leu Pro Pro Ile
25 30	35
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Pro Phe Pro Arg Phe Pro Trp Phe Arg Arg Asn	Phe Pro Ile Pro Ile
40 45 50	55
cct gaa tct gcc cct aca act ccc ctt cct agc	gaa aag taaacaaraa 354
Pro Glu Ser Ala Pro Thr Thr Pro Leu Pro Ser	Glu Lys
60 65	
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Lys Lys Val Leu Leu Ile Thr Ala Ile Leu Ala Val Ala Val Gly
   -15
                        -10
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Phe Pro Val Ser Gln Asp Xaa Glu Arg Glu Lys Arg Ser Ile Ser Asp
                                    10
age gat gaa tta get tea ggg ttt ttt gtg tte eet tae eea tat eea
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Ser Asp Glu Leu Ala Ser Gly Phe Phe Val Phe Pro Tyr Pro Tyr Pro
ttt cgc cca ctt cca cca att cca ttt cca aga ttt cca tgg ttt aga
                                                                      251
Phe Arg Pro Leu Pro Pro Ile Pro Phe Pro Arg Phe Pro Trp Phe Arg
                            40
egt aat ttt eet att eea ata eet gaa tet gee eet aca aet eee ett
                                                                      299
Arg Asn Phe Pro Ile Pro Ile Pro Glu Ser Ala Pro Thr Thr Pro Leu
                        55
                                            60
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                                                                      351
Pro Ser Glu Lys
65
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	11-
Met Thr Cys Arg Gly Ser -25	
tgc agc tac gct acc agg aga tct cca agc gaa ctc agc ctc cca	161
Cys Ser Tyr Ala Thr Arg Arg Ser Pro Ser Glu Leu Ser Leu Leu Pro	
-20 -15 -10	
ago too otg tgg gto ota goo aca ago tot oca aca att act att gca	209
Ser Ser Leu Trp Val Leu Ala Thr Ser Ser Pro Thr Ile Thr Ile Ala	
-5 1 5	
ctc gcg atg gcc gcc ggg aat ctg tgc ccc ctt cca tca tca tkt cgt	257
Leu Ala Met Ala Ala Gly Asn Leu Cys Pro Leu Pro Ser Ser Xaa Arg	
10 15 20 25	
crc aaa agg cgc tgg tgt cag gca asc car caa ara gct ctg ctg	302
	302
Xaa Lys Arg Arg Trp Cys Gln Ala Xaa Gln Gln Xaa Ala Leu Leu	
30 35 40	
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aaacaggctg ctggcattga ggtctgctac aaaaanarta atg gtc cca tgg ccc	175
Met Val Pro Trp Pro	2.4
•	
-55	
agg ggc aag gtg aaa act gct cct att ccc atc tct agg ttt cct ttc	223
Arg Gly Lys Val Lys Thr Ala Pro Ile Pro Ile Ser Arg Phe Pro Phe	
-50 -45 -40	
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Leu Pro Thr His Asp Pro Pro Thr Pro Ala His Trp Ser Pro Ala Ser	
-35 -30 -25 -20	
cat cag cag tit aaa cat kkg tca ccc ctc ctc act tig gcc cig cig	319
	213
His Gln Gln Phe Lys His Xaa Ser Pro Leu Leu Thr Leu Ala Leu Leu	
-15 -10 -5	
ggt cag tgc tct ctg ttc arc aat ttg agg aaa aaa ctt gca ggg caa	367
Gly Gln Cys Ser Leu Phe Xaa Asn Leu Arg Lys Lys Leu Ala Gly Gln	
1 5 10	
1 5 10 aaa gca aaa aaa tta cct tcc ttc tcc agc ctg ccc ctg aca ctc tgg	415

WO 99/31236 -209- PCT/IB98/02122

Lys Ala Lys Lys Leu Pro Ser Phe Ser Ser Leu Pro Leu Thr Leu Trp 15 20 25														
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Pro Leu Thr Pro Gln Phe Ala Glu Leu Thr Thr Val Ala Gln Lys Lys														
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Leu Val Pro Ala His Leu Ser Gly Leu Ile Thr Cys Leu Leu Ala Phe -20 -15 -10														
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Phe Gly Val Cys Pro Leu Leu Ser Val Thr Arg Val Val Ala Thr Glu
                -25
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                                                                      328
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                                -5
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Tyr Ile Ser Val Phe Cys Lys Val Thr Leu Ile
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att ctg acc ctc att ggc tgc ctg gtc aca ggc Ile Leu Thr Leu Ile Gly Cys Leu Val Thr Gly -10 -5												
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gag ggg aga gac ctg tcc gas tgg aaa aaa ggc : Glu Gly Arg Asp Leu Ser Xaa Trp Lys Lys Gly ( 115 120 125												
taaactggaa ctggacccag gatgctttgc ascaacgccc	tagggtttgc agtgaatgtc 691											
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							cca									222	2
Asn	Glu	Thr	Ile	Ile	Val	Leu	Pro -55	Ser	Asn	Val	Ile	Asn -50	Phe	Ser	Gln		
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Ala	Glu -45	Lys	Pro	Glu	Pro	Thr -40	Asn	Gln	Gly	Gln	Asp -35	Ser	Leu	Lys	Lys		
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	Leu	His	Ala	Glu		Lys	Val	Ile	Gly		Ile	Gln	Ile	Leu			
-30					-25					-20					-15	26	_
							999									360	•
GTA	Met	Met	vai	-10	Ser	neu	Gly	TIE	-5	nea	AIA	261	АІА	1	FIIE		
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	,	5					10					15			-		
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Pro		Ile	Gly	Pro	Phe		Phe	Ile	Ile	Ser		Ser	Leu	Ser	Ile		
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							aac									510	ט
	Thr	ьуs	ьуѕ	Arg	Leu 40	unr	Asn	Leu	Leu	va1 45	HIS	Inr	Thr	Leu	vai 50		
35	200	2++	ata	-a+		cta	tct	acc	cta		aat	ttc	att	avc		558	R
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Xaa	Lys	Asn 85	Asn	Ile	Pro	Thr	Xaa 90	Xaa	Tyr	Val	Xaa	Tyr 95	Phe	Tyr	His		
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Asp	Ser 100	Leu	Tyr	Thr	Thr	Asp 105	Xaa	Tyr	Thr	Ala	Lys 110	Ala	Xaa	Leu	Ala		
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Thr Gln Val Xaa Asp Thr Phe Ile Gly Tyr Arg Asn Leu Gly Phe Thr
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                          -20
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Ser Met Cys Ile Leu Phe His Cys Leu Leu Ser Phe Gln Val Phe Lys
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titing at gting ged cting and cting att cting gtt con the titing attention get the titing attention of the titing attention and the titing attention attention and the titing attention and the titing attention and the titing attention attention and the titing attention at	170
Met Val Ala Leu Asn Leu Ile Leu Val Pro Cys Cys Ala Ala Trp -10 -5	170
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5 10 15	
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Trp Arg Arg Ser Gln Pro Phe Ser Val Gly Leu Pro Ser Ala Glu Arg	
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Leu Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg Ser Leu Val Gly Glu	
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ggc gag cat ccs Gly Glu His Pro 100	rva btg tgaatkkkga Xaa Xaa 105	_ <del>_</del>	tga 441
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Val Ser Ser Trp 20	acg gag tgc ccg ccc Thr Glu Cys Pro Pro 25	Thr Trp Cys Ser Pr	o Leu Asp 35
Gln Val Cys Ile	tcc aac gag gtg gtc Ser Asn Glu Val Val 40	Val Ser Phe Lys Tr 45	p Ser Val 50
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cat	ttc	tgg	gat	. gg	aag	gga	tg:	t ga	g at	g ato	tgt	tao	tgo	aac Asr	ttc n Phe	304
Hls		Trp 40	ASŢ	, GT.	ν. ηλε	י פד)	45	5 GI	u Me	- TT6	_ cy:	50	y .			
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Lys	Arg 55	Ile	Ala	a Lev	ı Lev	Pro 60	Ly:	s Ar	g Ar	g Pho	e Let 65	ı Tr	o Th:	r Ly:	s Asp	
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400

Gly Glu His His Ser  90  tctagtttct atatagtgca atagagcata gattctataa attcttactt gtctaagaaa gtaaatctgt gttaaacaag tagtaataaa agttaattca atccaaaaaa aaaaaa 515  4210> 302  4211> 612  4212> DNA  4213> Homo sapiens  4220>  4221> S6268  4221> sig_peptide  4222> 56100  4223> Von Heijne matrix	Leu Phe Arg Asp Ser Leu Gln Gln Ser Met Arg Ile Phe Met Tyr Ser 70 75 80 85	'
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ctaatcgaaa agggggattt teeggtteeg geetggegag agtttgtgeg gegac atg  Met  —————————————————————————————————	· · · · · · · · · · · · · · · · · · ·	
Lys Leu Leu Thr His Asn Leu Leu Ser Ser His Val Arg Gly Val Gly  -10  tcc cgt ggc ttc ccc ctg cgc ctc cag gcc acc gag gtc cgt atc tgc  Ser Arg Gly Phe Pro Leu Arg Leu Gln Ala Thr Glu Val Arg Ile Cys  5  10  cct gtg gaa ttc aac ccc aac ttc gtg gcg cgt atg ata cct aaa gtg  Pro Val Glu Phe Asn Pro Asn Phe Val Ala Arg Met Ile Pro Lys Val  20  25  30  gag tgg tcg gcg ttc ctg gag gcg rmc gat aac ttg cgt ctg atc cag  Glu Trp Ser Ala Phe Leu Glu Ala Xaa Asp Asn Leu Arg Leu Ile Gln  35  40  45  50  gtg ccg aga agg gcc ggt tgagggatat gaggagaatg aggagttct  Val Pro Arg Arg Ala Gly  55  gaggaccatg caccacctgc tgctggaggt ggamstgaka gagggcaccc tgcagtgccc  ggaatctgga cgtatgttcc ccatcagccg cgggatcccc aacatgctgc tgagtgaaga  418  ggaaactgag agttgattgt gccaggcgc agtttttctt gttatgactg tgtatttttg  ttgatctata ccctqtttcc gaattctgcc gtgtgtatcc ccaacccttg acccaatgac  538	ctaatcgaaa agggggattt teeggtteeg geetggegag agtttgtgeg gegae atg Met	58
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Leu L	ieu i	ьуs	Val	-10	ьец	Leu	PIO	ren	-5	Pro	Ala	Ala	Ala	GIN 1	Asp		
tcg a Ser T																	149
	5	5					10					15			_		
caa c																	197
Gln A	119 1	PIIE	Pne	ALA	ьеи	25	1111	PIO	1111	пр	30	Ala	GIU	inr	inr		
tgc c																	245
Cys A	πā i	Leu	Arg	Ala	40	HIS	GIY	Cys	Arg	Asn 45	Pro	Thr	ьeu	vai	50		
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Ser A	sn I	Leu	Pro 70	Tyr	Ala	Ser	Trp	Phe 75	Glu	Ser	Phe	Сув	Gln 80	Phe	Thr		
cac t																	389
His T		Arg 85	Cys	ser	Asn	HIS	va1 90	Tyr	Tyr	Ala	Lys	Arg 95	Val	Leu	Cys		
tcc c																	437
Ser G	.00 :TD 1	Pro	Val	Ser	He	Leu 105	ser	Pro	Asn	Thr	Leu 110	Lys	Glu	Ile	Glu		
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															ctcat	1	887
tcgat	cago	cc c	agga	aata	ıg at	gaaa	itgaa	a tga	aaata	itat	gate	gagaa	ict o	cctac	tggag		947
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cgt ctt cag gaa gcc aga cag att caa gta ttg aag atg ctt cca agg Arg Leu Gln Glu Ala Arg Gln Ile Gln Val Leu Lys Met Leu Pro Arg	339
gaa aaa tta aga aga gaa gag aga aaa caa ata aat ggg aaa aaa Glu Lys Leu Arg Arg Glu Glu Arg Lys Gln Ile Asn Gly Lys Lys	387
raa agg aca aaa tat gaa aca cca aga aaa rga raa gga aaa aaa gga Xaa Arg Thr Lys Tyr Glu Thr Pro Arg Lys Xaa Xaa Gly Lys Lys Gly 30 35 40	435
gga aac mac cmc wtw tkt cmc ctt tcc aar agg gac tgaaactggg Gly Asn Xaa Xaa Xaa Xaa Leu Ser Lys Arg Asp 45 50 55	481
ctgacccttt tgatttccaa vctcascgtt ttggtgtaag gcggccaaar aaggatgcgg ascccagcac tgtgaagcct acaaaaacat tgatgcgctg gcttggggat ttgaatttga acatctttca cactaagttc agactcatga aaccaatctt cagatgctct gtaaaccaca taataaagag tttggaaatt aaaaaaaaar aa	541 601 661 693
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tgt Cys	ccg Pro	cgt Arg	caa Gln	gca Ala	acg Thr	cgc Arg	atc Ile	ccg Pro	ctc Leu	aac Asn	ggc Glv	acc Thr	tgg Trp	ctc Leu	ttc Phe	205
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Thr	Pro	Val	Ser 15	Lys	Met	Ala	Thr	Val 20	Lys	Ser	Glu	Leu	Ile 25	Glu	Arg	233
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		30					35	His		-		40			-	
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aac atg ccc acc act ggc ccc aac agc ctg agt tat gct agc tct gcc Asn Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala -20 -15 -10	277
ctg tcc ccc tgt ctg acc gct cca aag tcc ccc cga ctt gct atg atg Leu Ser Pro Cys Leu Thr Ala Pro Lys Ser Pro Arg Leu Ala Met Met -5 1 5 10	325
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Met Thr Pro Trp Cys Leu Ala Cys Leu Gly Arg Arg	
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ro Leu Ala Ser Leu Gln Trp Ser Leu Thr Leu Ala Trp Cys Gly Ser -10 -5 1	
go ago cao tgg aca gag aga coa akt cag akt toa cog tgg akt tot	326
Ser His Trp Thr Glu Arg Pro Xaa Gln Xaa Ser Pro Trp Xaa Ser 5 10 15	
tg tca gcg acc acc agg ggg tgatcacacg gaaggtgaac atccaggtcg	377
eu Ser Ala Thr Thr Arg Gly 20 25	
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cegeggtat ekteacagtg atcegggage tggaetaega taccaeremg geetaecage eweggtewa egecacagat caagacaara eeaggeetet gteeacestg geeaacttgg	617 677
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-20 -15 -10	
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Val Ile Pro Ser Ala Ala Ala Pro Ile His Asp Ala Asp Ala Gln Glu -5 1 5 10	
age tee ttg ggt ete aca gge ete cag age eta ete caa gge tte age	257
Ser Ser Leu Gly Leu Thr Gly Leu Gln Ser Leu Leu Gln Gly Phe Ser	
15 20 25	305
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					_	_		aac Asn					_			401
_		_	_					aag Lys								449
				-				cca Pro	_	-						497
_	_	Lys	_			_		gar Glu 115	_		_	_	_			545
	-	-	-	_	-			aca Thr	_						_	593
				_	_			cgg Arg				_	-	_	_	641
							-	aag Lys	_		_	_	-	-		689
				_				cac His								737
	_	_						ctg Leu 195			-			_		785
_				-				cag Gln	_	targ	gggt	ggg (	gacc	3999	ar	835
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agg acc ctg ctg gtt gct gcc ctc agg gcc tgg atg att cag tgc tgg Arg Thr Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln Cys Trp -10 -5 1 5	249
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ttr ggg gtc tac gtc atc cag gag cag gcg gcg gtc aag ctc cag tcc Leu Gly Val Tyr Val Ile Gln Glu Gln Ala Ala Val Lys Leu Gln Ser 25 30 35	345
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WO 99/31236

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														Met	Asn -60		
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ccc c Pro H	at					tct									cct	417	
tgt g Cys A		Thr		ccc		_	tgaa	accao		gtct	ccta	at co	ettt		a	468	
gcttc	aat	ga a gy c ca g	agga aago gtgc	cttt agag	t co	cttca ggaaa	aaaa atgga	c tgt a act	agco	ctcc	tctc	acto	gaa g ggt a	ggtgg	accata ggagct acagag	528 588 648 697	
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cta c	ctc	acc	tgg	ctt	ttc	aca	cta			ttg	atc	atg			ttg	222	
Leu I	Leu	Thr -15	Trp	Leu	Phe	Thr	Leu -10	Leu	Phe	Leu	Ile	Met -5	Leu	Val	Leu	<i>-</i> -	
Lys I																270	1
gtc t		ata	ttt	gat		atc	ctt	ctt	gtc		ctg	att	gtg	aaa		318	ļ
Val 1																	

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Met	1,,
atc cct ctg ata age cac ctt gcc gag gct gct cct cct acc tca tgg	225
Ile Pro Leu Ile Ser His Leu Ala Glu Ala Ala Pro Pro Thr Ser Trp	223
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Ser Leu Ile Ser Ser Val Leu Asn Val Gly His Leu Leu Phe Ser Ser	2,5
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Ala Cys Ser Val Ser Leu Glu Ala Leu Ser Thr Arg Asn Ile Lys Ala	321
1 5 10 15	
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ile ile bed met bys	
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	436 496
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           Met Glu Ser Pro Gln Leu His Cys Ile Leu Asn Ser Asn
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Ser Val Ala Cys Ser Phe Ala Val Gly Ala Gly Phe Leu Ala Phe Leu
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-25
                    -20
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1112

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526

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                                          Met Thr Pro Arg Ile Leu
age gaa gte cag ttt tea gea ttt tgt eet tat tgg aca ata gea agg
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                                                -30
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                        -20
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Val Val Cys Val Ile Cys Val Thr Leu Asn Phe Pro Arg Phe Tyr Phe
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Leu Cys Leu Ser Ser Leu Thr Ala Phe Gly Thr Pro Pro Ile Gly Val
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cac att ccc tct ccc tararcacac tcccttggat ttcctcradt ggggtctgct
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Phe Thr Ile Lys Ser Leu Ser Arg Pro Gly Gln Pro Trp Cys Glu Ala
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His Val Phe Leu Asn Lys Asn Leu Phe Leu Gln Tyr Asn Ser Asp Asn
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Pro Ser Thr Leu Gln Val Xaa Xaa Phe Cys Gln Arg Glu Ala Glu Arg
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Cys Thr Gly Ala Ser Trp Gln Phe Ala Thr Asn Gly Glu Lys Ser Leu
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Ser Xaa Ile Lys Glu Thr Trp Lys Lys Asp Arg Xaa Leu Glu Xaa Tyr
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                                                                      631
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 Phe Arg Lys Leu Ser Lys Gly Asp Cys Asp His Trp Leu Arg Glu Phe
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				_	cc aa		_							5000	-090	989
-	-900.		Juan			-5					~~~	••				,,,,
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687

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